

# WORLD INTELLECTUAL PROPERTY ORGANIZATION



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60/098,639 60/117,393 31 August 1998 (31.08.98) 27 January 1999 (27.01.99) (US). MONAHAN, John, E. [US/US]; 942 West Street, Walpole, MA 02081 (US). SCHLEGEL, Robert [US/US]; 211 Melrose Street, Auburndale, MA 02466 (US).

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(54) Title: HUMAN GENES DIFFERENTIALLY EXPRESSED IN COLORECTAL CANCER

(57) Abstract

This invention relates to novel human genes, to proteins expressed by the genes, and to variants of the proteins. The invention also relates to diagnostic assays and therapeutic agents related to the genes and proteins, including probes, antisense constructs, and antihodies. The subject nucleic acids have been found to be differentially regulated in tumor cells, particularly in colon cancer tissue.

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Int Jonal Application No PCT/US 99/19424

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Category *	Citation of document, with indication, where appropriate, of the re	alovant passages		Palauant to alaim No
		prevant passages		Relevant to claim No.
Х	EMBL Database Hum2:SEQ ID HS23K2 Human DNA sequence from clone 23 chromosome Xq25-26.2 XP002126133	3K20 on		2,8,15,
	compare nucleotides 37007-37027 with nucleotides 513-493 in SEQ			
X	EMBL Database Hum2:SEQ ID HS24M1 Human DNA sequence from PAC 24M1 chromosome 1, contains tenascin- (restrictin). 16 April 1997 XP002126134	15 on -R		8,15,30
:	compare nucleotides 98498-98510 with nucleotides 506-494 in SEQ this application			
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X Furti	her documents are listed in the continuation of box C.	X Patent fami	ly members as	re listed in annex.
° Special ca	tegories of cited documents :	Tre lates de	ushlish s = 4 - 4	Ab. (1.4 1.47) - 1.47(1)
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	7 December 1999	Pare or maning (		onal search report 5. <b>04</b> , 2000
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	NL - 2280 HV Rijawijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	CUPIDO	), M	

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C.(Continua Category °	tion) DOCUMENTS CONSIDERED TO BE RELEVANT  Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A		1
	SCHWEINFEST C W ET AL: "Subtraction hybridization cDNA libraries from colon carcinoma and hepatic cancer" GENE ANALYSIS TECHNIQUES, vol. 7, 1 January 1990 (1990-01-01), pages 64-70, XP002089887 ISSN: 0735-0651 page 64	1,18
A	VIDER B ET AL: "Human colorectal carcinogenesis is associated with deregulation of homeobox gene expression" BIOCHEMICAL AND BIOPHYSICAL RESEARCH COMMUNICATIONS, vol. 232, no. 3, March 1997 (1997-03), pages 742-748, XP002104685 ISSN: 0006-291X page 742	1
A	JAU MIN WONG ET AL: "UBIQUITIN-RIBOSOMAL PROTEIN S27A GENE OVEREXPRESSES IN HUMAN COLORECTAL CARCINOMA IS AN EARLY GROWTH RESPONSE GENE" CANCER RESEARCH, vol. 53, no. 8, 15 April 1993 (1993-04-15), pages 1916-1920, XP002024627 ISSN: 0008-5472 page 1916	
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	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
tegory °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	WO 95 11923 A (DANA FARBER CANCER INST INC) 4 May 1995 (1995-05-04) page 1, line 29 -page 6, line 17 page 19, line 7 -page 29, line 11	1-6,9, 10,14, 17-25, 31-34
<b>\</b>	EP 0 284 362 A (ICI PLC) 28 September 1988 (1988-09-28) the whole document	1-25, 27-34

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international application No.

PCT/US 99/19424

Box i Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
2. X Claims Nos.: 26 because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
see FURTHER INFORMATION sheet PCT/ISA/210
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. X  No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
1-25,27,28,29-34 (all partially)
Remark on Protest  The additional search fees were accompanied by the applicant's protest.
No protest accompanied the payment of additional search fees.
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### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box 3.

Claims Nos.: 26

Claim 26 relates to an agent which alters the expression in a cell of a nucleic acid. As this agent has not been disclosed in the application, a meaningful search could not be performed.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

# FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Claims: 1-25, 27,28, 29-34, all partially

Invention 1:

An isolated nucleic acid comprising a nucleotide sequence which hybridizes under stringent conditions to SEQ.ID.No:1 or a sequence complementary thereto; an isolated nucleic acid, comprising a nucleotide sequence at least 80% identical to at least 15 consecutive nucleotides of SEQ.ID. No:1 or a sequence complementary thereto; an isolated nucleic acid comprising nucleotide sequence of SEQ.ID.No:1 or a sequence complementary thereto; an expression vector comprising said nucleic acids; an host cell comprising said vector; a transgenic animal having a transgene comprising said nucleic acids; a nucleic acid hybridizing to a nucleic acid probe corresponding to at least 12 consecutive nucleotides of SEQ.ID.No:1; a probe/primer hybridizing to a nucleic acid probe corresponding to at least 12 consecutive nucleotides of SEQ.ID.No:1; an isolated polypeptide encoded by said nucleic acid; an antibody that specifically binds to said polypeptide; an antisense oligonucleotide which hybridizes under stringent conditions to at least 12 consecutive nucleic acids of SEQ.ID.No:1; a test kit comprising said probe/primer; a testkit comprising said antiboda; a method for determining the phenotype of a cell comprising detecting the differential expression of a nucleic acid which hybridizes under stringent conditions to at least 12 consecutive nucleic acids of SEQ.ID. No:1 or a protein encoded by said nucleic acid; a method for determing the presence or absence of a nucleic acid which hybridizes under stringent conditions to at least 12 consecutive nucleic acids of SEQ.ID.No:1; a method for detecting a mutation in a test nucleic acid which hybridizes under stringent conditions to at least 12 consecutive nucleic acids of SEQ.ID.No:1; a method for identifying an agent which alters the level of expression in a cell of a nucleic acid which hybridizes under stringent conditions to at least 12 consecutive nucleic acids of SEQ.ID.No:1; a pharmaceutical composition comprising a nucleic acid which hybridizes under stringent conditions to at least 12 consecutive nucleic acids of SEQ.ID.No:1; a pharmaceutical composition comprising a polypeptide encoded by said nucleic acid; a method for detecting cancer using SEQ.ID.No:1 or an antibody to a protein encoded by said sequence, as a probe.

Claims: 1-25, 27,28 30-34, all partially

Inventions 2 to 35:

Idem as invention 1, wherein each invention relates to the nucleic acid encoded by SEQ.ID.Nos:2 to 35 in stead of SEQ.ID.No:1.

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Claims: 8,10-14,16-22,24,25,27,29,31-34, all partially

Inventions 36 to 168:

Idem as invention 1, wherein each invention relates to the nucleic acid encoded by SEQ.ID.Nos:36 to 168 in stead of SEQ.ID.No:1.

Claims: 16-21,24,25, 27,28,31-34, all partially

Inventions 169 to 544:

Idem as invention 1, wherein each invention relates to the nucleic acid encoded by SEQ.ID.Nos:169 to 544 in stead of SEQ.ID.No:1.

page 2 of 2

Information on patent family members

Int. .tional Application No PCT/US 99/19424

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## **PCT**

# WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



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(75) Inventors/Applicants (for US only): ENDEGE, Wilson, O. [KE/US]; 222 Normandy Drive, Norwood, MA 02062 (US). STEINMANN, Kathleen, E. [US/US]; 115 Washington Street, Unit 3B, Winchester, MA 01890 (US). ASTLE, Jon, H. [US/US]; 42 Short Street, Taunton, MA 02780 (US). BURGESS, Christopher, C. [US/US]; 97 Canton Terrace, Westwood, MA 02090 (US). CARROLL, Eddie, III [US/US]; Apartment 3, 1175 Washington Street, Norwood, MA 02062 (US). CATINO, Theodore, J. [US/US]; 18 Jo Paul Drive, Attleboro, MA 02702 (US). DWIVEDI, Poornima [US/US]; 10 Haven Road, Medfield, MA 02052 (US). FORD, Donna, M. [US/US]; 8 Morningside Road,

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#### Published

NE, SN, TD, TG).

Without international search report and to be republished upon receipt of that report.

(54) Title: HUMAN GENES DIFFERENTIALLY EXPRESSED IN COLORECTAL CANCER

#### (57) Abstract

This invention relates to novel human genes, to proteins expressed by the genes, and to variants of the proteins. The invention also relates to diagnostic assays and therapeutic agents related to the genes and proteins, including probes, antisense constructs, and antibodies. The subject nucleic acids have been found to be differentially regulated in tumor cells, particularly in colon cancer tissue.

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# HUMAN GENES DIFFERENTIALLY EXPRESSED IN COLORECTAL CANCER

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#### **Related Application Information**

This application is based on Provisional Application Nos. 60/117,393, filed January 27, 1999, and 60/098,639, filed August 31, 1998, which are incorporated herein by reference in their entirety.

#### Field of the Invention

The present invention provides nucleic acid sequences and proteins encoded thereby, as well as probes derived from the nucleic acid sequences, antibodies directed to the encoded proteins, and diagnostic methods for detecting cancerous cells, especially colon cancer cells.

## Background of the Invention

Colorectal carcinoma is a malignant neoplastic disease. There is a high incidence of colorectal carcinoma in the Western world, particularly in the United States. Tumors of this type often metastasize through lymphatic and vascular channels. Many patients with colorectal carcinoma eventually die from this disease. In fact, it is estimated that 62,000 persons in the United States alone die of colorectal carcinoma annually.

However, if diagnosed early, colon cancer may be treated effectively by surgical removal of the cancerous tissue. Colorectal cancers originate in the colorectal epithelium and typically are not extensively vascularized (and therefore not invasive) during the early stages of development. Colorectal cancer is thought to result from the clonal expansion of a single mutant cell in the epithelial lining of the colon or rectum. The transition to a highly vascularized, invasive and ultimately metastatic cancer which spreads throughout the body commonly takes ten years or longer. If the cancer is detected prior to invasion, surgical removal of the cancerous tissue is an effective cure. However, colorectal cancer is often detected only upon manifestation of clinical symptoms, such as pain and black tarry stool. Generally, such symptoms are present only when the disease is well established, often after metastasis has occurred, and the

prognosis for the patient is poor, even after surgical resection of the cancerous tissue. Early detection of colorectal cancer therefore is important in that detection may significantly reduce its morbidity.

Invasive diagnostic methods such as endoscopic examination allow for direct visual identification, removal, and biopsy of potentially cancerous growths such as polyps. Endoscopy is expensive, uncomfortable, inherently risky, and therefore not a practical tool for screening populations to identify those with colorectal cancer. Non-invasive analysis of stool samples for characteristics indicative of the presence of colorectal cancer or precancer is a preferred alternative for early diagnosis, but no known diagnostic method is available which reliably achieves this goal. A reliable, non-invasive, and accurate technique for diagnosing colon cancer at an early stage would help save many lives.

#### Summary of the Invention

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The present invention provides nucleic acid sequences and proteins encoded thereby, as well as probes derived from the nucleic acid sequences, antibodies directed to the encoded proteins, and diagnostic methods for detecting cancerous cells, especially colon cancer cells. The sequences disclosed herein have been found to be differentially expressed in samples obtained from colon cancer cell lines and/or colon cancer tissue. The 544 sequences that were obtained were analyzed by "blasting" the sequences against the publicly available databases; based upon the Blast search results it was found that SEQ ID Nos: 1-35 contained novel sequences, SEQ ID Nos: 36-168 contained EST sequences and SEQ ID Nos: 169-544 contained known sequences.

In one aspect, the invention provides an isolated nucleic acid comprising a nucleotide sequence which hybridizes under stringent conditions to a sequence of SEQ ID Nos. 1-544 or a sequence complementary thereto. In a related embodiment, the nucleic acid is at least about 80% or about 100% identical to a sequence corresponding to at least about 12, at least about 15, at least about 25, or at least about 40 consecutive nucleotides up to the full length of one of SEQ ID Nos. 1-544 or a sequence complementary thereto or up to the full length of the gene of which said sequence is a fragment. In certain embodiments, a nucleic acid of the present invention includes at least about five, at least about ten, or at least about twenty

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nucleic acids from a region designated as novel in Table 2. In certain other embodiments, a nucleic acid of the present invention includes at least about five, at least about ten, or at least about twenty nucleotides which are not included in corresponding clones whose accession numbers are listed in Table 2.

In another aspect, the invention provides an isolated nucleic acid comprising a nucleotide sequence which hybridizes under stringent conditions to a sequence of SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35, or a sequence complementary thereto. In a related embodiment, the nucleic acid is at least about 80% or about 100% identical to a sequence corresponding to at least about 12, at least about 15, at least about 25, or at least about 40 consecutive nucleotides up to the full length of one of SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35 or a sequence complementary thereto or up to the full length of the gene of which said sequence is a fragment. In certain embodiments, a nucleic acid of the present invention includes at least about five, at least about ten, or at least about twenty nucleic acids from a region designated as novel in Table 2. In certain other embodiments, a nucleic acid of the present invention includes at least about five, at least about ten, or at least about twenty nucleic acid of the present invention includes at least about five, at least about ten, or at least about twenty nucleotides which are not included in corresponding clones whose accession numbers are listed in Table 2.

In one embodiment, the invention provides a nucleic acid comprising a nucleotide sequence which hybridizes under stringent conditions to a sequence of SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, and a transcriptional regulatory sequence operably linked to the nucleotide sequence to render the nucleotide sequence suitable for use as an expression vector. In another embodiment, the nucleic acid may be included in an expression vector capable of replicating in a prokaryotic or eukaryotic cell. In a related embodiment, the invention provides a host cell transfected with the expression vector.

In another embodiment, the invention provides a transgenic animal having a transgene of a nucleic acid comprising a nucleotide sequence which hybridizes under stringent conditions to a sequence of SEQ ID Nos. 1-168, preferably SEQ ID Nos 1-35, or a sequence complementary thereto incorporated in cells thereof. The transgene modifies the level of expression of the nucleic acid, the stability of a mRNA transcript of the nucleic acid, or the activity of the encoded product of the nucleic acid.

In yet another embodiment, the invention provides substantially pure nucleic acid which hybridizes under stringent conditions to a nucleic acid probe corresponding to at least about 12, at least about 15, at least about 25, or at least about 40 consecutive nucleotides up to the full length of one of SEQ ID Nos. 1-168, preferably SEQ ID Nos 1-35, or a sequence complementary thereto or up to the full length of the gene of which said sequence is a fragment. The invention also provides an antisense oligonucleotide analog which hybridizes under stringent conditions to at least 12, at least 25, or at least 50 consecutive nucleotides of one of SEQ ID Nos. 1-544 up to the full length of one of SEQ ID Nos. 1-544 or a sequence complementary thereto or up to the full length of the gene of which said sequence is a fragment, and which is resistant to cleavage by a nuclease, preferably an endogenous endonuclease or exonuclease.

In another embodiment, the invention provides a probe/primer comprising a substantially purified oligonucleotide, said oligonucleotide containing a region of nucleotide sequence which hybridizes under stringent conditions to at least about 12, at least about 15, at least about 25, or at least about 40 consecutive nucleotides of sense or antisense sequence selected from SEQ ID Nos. 1-168 up to the full length of one of SEQ ID Nos. 1-168 or a sequence complementary thereto or up to the full length of the gene of which said sequence is a fragment. In preferred embodiments, the probe selectively hybridizes with a target nucleic acid. In another embodiment, the probe may include a label group attached thereto and able to be detected. The label group may be selected from radioisotopes, fluorescent compounds, enzymes, and enzyme co-factors. The invention further provides arrays of at least about 10, at least about 25, at least about 50, or at least about 100 different probes as described above attached to a solid support.

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In yet another embodiment, the invention pertains to a method of determining the phenotype of a cell, comprising detecting the differential expression, relative to a normal cell, of at least one nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-544, wherein the nucleic acid is differentially expressed by at least a factor of two, at least a factor of five, at least a factor of twenty, or at least a factor of fifty.

In another aspect, the invention provides polypeptides encoded by the subject nucleic acids. In one embodiment, the invention pertains to a polypeptide including an

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amino acid sequence encoded by a nucleic acid comprising a nucleotide sequence which hybridizes under stringent conditions to a sequence of SEQ ID Nos. 1-168 or a sequence complementary thereto, or a fragment comprising at least about 25, or at least about 40 amino acids thereof. Further provided are antibodies immunoreactive with these polypeptides.

In still another aspect, the invention provides diagnostic methods. In one embodiment, the invention pertains to a method for determining the phenotype of cells from a patient by providing a nucleic acid probe comprising a nucleotide sequence having at least 12, at least about 15, at least about 25, or at least about 40 consecutive nucleotides represented in a sequence of SEQ ID Nos. 1-544 up to the full length of one of SEQ ID Nos. 1-544 or a sequence complementary thereto or up to the full length of the gene of which said sequence is a fragment, obtaining a sample of cells from a patient, providing a second sample of cells substantially all of which are non-cancerous, contacting the nucleic acid probe under stringent conditions with mRNA of each of said first and second cell samples, and comparing (a) the amount of hybridization of the probe with mRNA of the first cell sample, with (b) the amount of hybridization of the probe with mRNA of the second cell sample, wherein a difference of at least a factor of two, at least a factor of five, at least a factor of twenty, or at least a factor of fifty in the amount of hybridization with the mRNA of the first cell sample as compared to the amount of hybridization with the mRNA of the second cell sample is indicative of the phenotype of cells in the first cell sample. Determining the phenotype includes determining the genotype, as the term is used herein.

In another embodiment, the invention provides a test kit for identifying an transformed cells, comprising a probe/primer as described above, for measuring a level of a nucleic acid which hybridizes under stringent conditions to a nucleic acid of SEQ ID Nos. 1-544 in a sample of cells isolated from a patient. In certain embodiments, the kit may further include instructions for using the kit, solutions for suspending or fixing the cells, detectable tags or labels, solutions for rendering a nucleic acid susceptible to hybridization, solutions for lysing cells, or solutions for the purification of nucleic acids.

In another embodiment, the invention provides a method of determining the phenotype of a cell, comprising detecting the differential expression, relative to a normal cell, of at least one protein encoded by a nucleic acid which hybridizes under

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stringent conditions to one of SEQ ID Nos. 1-544, wherein the protein is differentially expressed by at least a factor of two, at least a factor of five, at least a factor of twenty, or at least a factor of fifty. In one embodiment, the level of the protein is detected in an immunoassay. The invention also pertains to a method for determining the presence or absence of a nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-168 in a cell, comprising contacting the cell-with a probe as described above. The invention further provides a method for determining the presence or absence of a subject polypeptide encoded by a nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-168 in a cell, comprising contacting the cell with an antibody as described above. In yet another embodiment, the invention provides a method for determining the presence of an aberrant mutation (e.g., deletion, insertion, or substitution of nucleic acids) or aberrant methylation in a gene which hybridizes under stringent conditions to a sequence of SEQ ID Nos. 1-168 or a sequence complementary thereto, comprising collecting a sample of cells from a patient, isolating nucleic acid from the cells of the sample. contacting the nucleic acid sample with one or more primers which specifically hybridize to a nucleic acid sequence of SEQ ID Nos. 1-544 under conditions such that hybridization and amplification of the nucleic acid occurs, and comparing the presence, absence, or size of an amplification product to the amplification product of a normal cell.

In one embodiment, the invention provides a test kit for identifying transformed cells, comprising an antibody specific for a protein encoded by a nucleic acid which hybridizes under stringent conditions to any one of SEQ Nos. 1-544. In certain embodiments, the kit further includes instructions for using the kit. In certain embodiments, the kit may further include instructions for using the kit, solutions for suspending or fixing the cells, detectable tags or labels, solutions for rendering a polypeptide susceptible to the binding of an antibody, solutions for lysing cells, or solutions for the purification of polypeptides.

In yet another aspect, the invention provides pharmaceutical compositions including the subject nucleic acids. In one embodiment, an agent which alters the level of expression in a cell of a nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-544 or a sequence complementary thereto is identified by providing a cell, treating the cell with a test agent, determining the level

of expression in the cell of a nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-544 or a sequence complementary thereto, and comparing the level of expression of the nucleic acid in the treated cell with the level of expression of the nucleic acid in an untreated cell, wherein a change in the level of expression of the nucleic acid in the treated cell relative to the level of expression of the nucleic acid in the untreated cell is indicative of an agent which alters the level of expression of the nucleic acid in a cell. The invention further provides a pharmaceutical composition comprising an agent identified by this method. In another embodiment, the invention provides a pharmaceutical composition which includes a polypeptide encoded by a nucleic acid having a nucleotide sequence that hybridizes under stringent conditions to one of SEQ ID Nos. 1-544 or a sequence complementary thereto. In one embodiment, the invention pertains to a pharmaceutical composition comprising a nucleic acid including a sequence which hybridizes under stringent conditions to one of SEQ ID Nos. 1-544 or a sequence complementary thereto.

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## Brief Description of the Figure

The figure depicts an exemplary assay result for determining differential expression of gene products in cells.

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## Detailed Description of the Invention

The invention relates to nucleic acids having the disclosed nucleotide sequences (SEQ ID Nos. 1-544), as well as full length cDNA, mRNA, and genes corresponding to these sequences, and to polypeptides and proteins encoded by these nucleic acids and genes, and portions thereof.

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Also included are polypeptides and proteins encoded by the nucleic acids of SEQ ID Nos. 1-544. The various nucleic acids that can encode these polypeptides and proteins differ because of the degeneracy of the genetic code, in that most amino acids are encoded by more than one triplet codon. The identity of such codons is well known in this art, and this information can be used for the construction of the nucleic acids within the scope of the invention.

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Nucleic acids encoding polypeptides and proteins that are variants of the polypeptides and proteins encoded by the nucleic acids and related cDNA and genes are also within the scope of the invention. The variants differ from wild-type protein in having one or more amino acid substitutions that either enhance, add, or diminish a

biological activity of the wild-type protein. Once the amino acid change is selected, a nucleic acid encoding that variant is constructed according to the invention.

The following detailed description discloses how to obtain or make full-length cDNA and human genes corresponding to the nucleic acids, how to express these nucleic acids and genes, how to identify structural motifs of the genes, how to identify the function of a protein encoded by a gene corresponding to an nucleic acid, how to use nucleic acids as probes in mapping and in tissue profiling, how to use the corresponding polypeptides and proteins to raise antibodies, and how to use the nucleic acids, polypeptides, and proteins for therapeutic and diagnostic purposes.

The sequences investigated herein have been found to be differentially expressed in samples obtained from colon cancer tissue. However, it is also believed that these sequences may also have utility with other types of cancer. In a related application, PCT/IB99/01062, filed June 9, 1999, the inventors disclosed nucleic acid sequences that are differentially expressed in colon cancer-derived cell lines, such as SW 480, relative to the expression levels in normal tissue, e.g., normal colon tissue and/or normal non-colon tissue. In this application, Table 3 lists nucleic acid sequences which are over-expressed in both cancer cell line SW 480 as well colon cancer tissue obtained from various patients.

Accordingly, certain aspects of the present invention relate to nucleic acids differentially expressed in tumor tissue, especially colon cancer cell lines, polypeptides encoded by such nucleic acids, and antibodies immunoreactive with these polypeptides, and preparations of such compositions. Moreover, the present invention provides diagnostic and therapeutic assays and reagents for detecting and treating disorders involving, for example, aberrant expression of the subject nucleic acids.

#### I. General

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This invention relates in part to novel methods for identifying and/or classifying cancerous cells present in a human tumors, particularly in solid tumors, e.g., carcinomas and sarcomas, such as, for example, breast or colon cancers. The method uses genes that are differentially expressed in cancer cell lines and/or cancer tissue compared with related normal cells, such as normal colon cells, and thereby

identifies or classifies tumor cells by the upregulation and/or downregulation of expression of particular genes, an event which is implicated in tumorigenesis.

Upregulation or increased expression of certain genes such as oncogenes, act to promote malignant growth. Downregulation or decreased expression of genes such as tumor suppressor genes also promotes malignant growth. Thus, alteration in the expression of either type of gene is a potential diagnostic indicator-for determining whether a subject is at risk of developing or has cancer, e.g., colon cancer.

Accordingly, in one aspect, the invention also provides biomarkers, such as nucleic acid markers, for human tumor cells, e.g., for colon cancer cells. The invention also provides proteins encoded by these nucleic acid markers.

The invention also features methods for identifying drugs useful for treatment of such cancer cells, and for treatment of a cancerous condition, such as colon cancer. Unlike prior methods, the invention provides a means for identifying cancer cells at an early stage of development, so that premalignant cells can be identified prior to their spreading throughout the human body. This allows early detection of potentially cancerous conditions, and treatment of those cancerous conditions prior to spread of the cancerous cells throughout the body, or prior to development of an irreversible cancerous condition.

## II. <u>Definitions</u>

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For convenience, the meaning of certain terms and phrases used in the specification, examples, and appended claims, are provided below.

The term "an aberrant expression", as applied to a nucleic acid of the present invention, refers to level of expression of that nucleic acid which differs from the level of expression of that nucleic acid in healthy tissue, or which differs from the activity of the polypeptide present in a healthy subject. An activity of a polypeptide can be aberrant because it is stronger than the activity of its native counterpart. Alternatively, an activity can be aberrant because it is weaker or absent relative to the activity of its native counterpart. An aberrant activity can also be a change in the activity; for example, an aberrant polypeptide can interact with a different target peptide. A cell can have an aberrant expression level of a gene due to overexpression or underexpression of that gene.

The term "agonist", as used herein, is meant to refer to an agent that mimics or upregulates (e.g., potentiates or supplements) the bioactivity of a protein. An agonist can be a wild-type protein or derivative thereof having at least one bioactivity of the wild-type protein. An agonist can also be a compound that upregulates expression of a gene or which increases at least one bioactivity of a protein. An agonist can also be a compound which increases the interaction of a polypeptide with another molecule, e.g., a target peptide or nucleic acid.

The term "allele", which is used interchangeably herein with "allelic variant", refers to alternative forms of a gene or portions thereof. Alleles occupy the same locus or position on homologous chromosomes. When a subject has two identical alleles of a gene, the subject is said to be homozygous for that gene or allele. When a subject has two different alleles of a gene, the subject is said to be heterozygous for the gene. Alleles of a specific gene can differ from each other in a single nucleotide, or several nucleotides, and can include substitutions, deletions, and/or insertions of nucleotides. An allele of a gene can also be a form of a gene containing mutations.

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The term "allelic variant of a polymorphic region of a gene" refers to a region of a gene having one of several nucleotide sequences found in that region of the gene in other individuals.

"Antagonist" as used herein is meant to refer to an agent that downregulates (e.g., suppresses or inhibits) at least one bioactivity of a protein. An antagonist can be a compound which inhibits or decreases the interaction between a protein and another molecule, e.g., a target peptide or enzyme substrate. An antagonist can also be a compound that downregulates expression of a gene or which reduces the amount of expressed protein present.

The term "antibody" as used herein is intended to include whole antibodies, e.g., of any isotype (IgG, IgA, IgM, IgE, etc), and includes fragments thereof which are also specifically reactive with a vertebrate, e.g., mammalian, protein. Antibodies can be fragmented using conventional techniques and the fragments screened for utility in the same manner as described above for whole antibodies. Thus, the term includes segments of proteolytically-cleaved or recombinantly-prepared portions of an antibody molecule that are capable of selectively reacting with a certain protein.

Nonlimiting examples of such proteolytic and/or recombinant fragments include Fab, F(ab')2, Fab', Fv, and single chain antibodies (scFv) containing a V[L] and/or V[H]

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domain joined by a peptide linker. The scFv's may be covalently or non-covalently linked to form antibodies having two or more binding sites. The subject invention includes polyclonal, monoclonal, or other purified preparations of antibodies and recombinant antibodies.

The phenomenon of "apoptosis" is well known, and can be described as a programmed death of cells. As is known, apoptosis is contrasted with "necrosis", a phenomenon when cells die as a result of being killed by a toxic material, or other external effect. Apoptosis involves chromatic condensation, membrane blebbing, and fragmentation of DNA, all of which are generally visible upon microscopic examination.

A disease, disorder, or condition "associated with" or "characterized by" an aberrant expression of a nucleic acid refers to a disease, disorder, or condition in a subject which is caused by, contributed to by, or causative of an aberrant level of expression of a nucleic acid.

As used herein the term "bioactive fragment of a polypeptide" refers to a fragment of a full-length polypeptide, wherein the fragment specifically agonizes (mimics) or antagonizes (inhibits) the activity of a wild-type polypeptide. The bioactive fragment preferably is a fragment capable of interacting with at least one other molecule, e.g., protein, small molecule, or DNA, which a full length protein can bind.

"Biological activity" or "bioactivity" or "activity" or "biological function", which are used interchangeably, herein mean an effector or antigenic function that is directly or indirectly performed by a polypeptide (whether in its native or denatured conformation), or by any subsequence thereof. Biological activities include binding to polypeptides, binding to other proteins or molecules, activity as a DNA binding protein, as a transcription regulator, ability to bind damaged DNA, etc. A bioactivity can be modulated by directly affecting the subject polypeptide. Alternatively, a bioactivity can be altered by modulating the level of the polypeptide, such as by modulating expression of the corresponding gene.

The term "biomarker" refers a biological molecule, e.g., a nucleic acid, peptide, hormone, etc., whose presence or concentration can be detected and correlated with a known condition, such as a disease state.

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"Cells," "host cells", or "recombinant host cells" are terms used interchangeably herein. It is understood that such terms refer not only to the particular subject cell but to the progeny or potential progeny of such a cell. Because certain modifications may occur in succeeding generations due to either mutation or environmental influences, such progeny may not, in fact, be identical to the parent cell, but are still included within the scope of the term as used herein.

A "chimeric polypeptide" or "fusion polypeptide" is a fusion of a first amino acid sequence encoding one of the subject polypeptides with a second amino acid sequence defining a domain (e.g., polypeptide portion) foreign to and not substantially homologous with any domain of the subject polypeptide. A chimeric polypeptide may present a foreign domain which is found (albeit in a different polypeptide) in an organism which also expresses the first polypeptide, or it may be an "interspecies," "intergenic," etc., fusion of polypeptide structures expressed by different kinds of organisms. In general, a fusion polypeptide can be represented by the general formula  $(X)_n-(Y)_m-(Z)_n$ , wherein Y represents a portion of the subject polypeptide, and X and Z are each independently absent or represent amino acid sequences which are not related to the native sequence found in an organism, or which are not found as a polypeptide chain contiguous with the subject sequence, where m is an integer greater than or equal to one, and each occurrence of n is, independently, 0 or an integer greater than or equal to 1 (n and m are preferably no greater than 5 or 10).

A "delivery complex" shall mean a targeting means (e.g., a molecule that results in higher affinity binding of a nucleic acid, protein, polypeptide or peptide to a target cell surface and/or increased cellular or nuclear uptake by a target cell). Examples of targeting means include: sterols (e.g., cholesterol), lipids (e.g., a cationic lipid, virosome or liposome), viruses (e.g., adenovirus, adeno-associated virus, and retrovirus), or target cell-specific binding agents (e.g., ligands recognized by target cell specific receptors). Preferred complexes are sufficiently stable *in vivo* to prevent significant uncoupling prior to internalization by the target cell. However, the complex is cleavable under appropriate conditions within the cell so that the nucleic acid, protein, polypeptide or peptide is released in a functional form.

As is well known, genes or a particular polypeptide may exist in single or multiple copies within the genome of an individual. Such duplicate genes may be identical or may have certain modifications, including nucleotide substitutions,

additions or deletions, which all still code for polypeptides having substantially the same activity. The term "DNA sequence encoding a polypeptide" may thus refer to one or more genes within a particular individual. Moreover, certain differences in nucleotide sequences may exist between individual organisms, which are called alleles. Such allelic differences may or may not result in differences in amino acid sequence of the encoded polypeptide yet still encode a polypeptide with the same biological activity.

The term "equivalent" is understood to include nucleotide sequences encoding functionally equivalent polypeptides. Equivalent nucleotide sequences will include sequences that differ by one or more nucleotide substitutions, additions or deletions, such as allelic variants; and will, therefore, include sequences that differ from the nucleotide sequence of the nucleic acids shown in SEQ ID NOs: 1-544 due to the degeneracy of the genetic code.

As used herein, the terms "gene", "recombinant gene", and "gene construct" refer to a nucleic acid of the present invention associated with an open reading frame, including both exon and (optionally) intron sequences.

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A "recombinant gene" refers to nucleic acid encoding a polypeptide and comprising exon sequences, though it may optionally include intron sequences which are derived from, for example, a related or unrelated chromosomal gene. The term "intron" refers to a DNA sequence present in a given gene which is not translated into protein and is generally found between exons.

The term "growth" or "growth state" of a cell refers to the proliferative state of a cell as well as to its differentiative state. Accordingly, the term refers to the phase of the cell cycle in which the cell is, e.g., G0, G1, G2, prophase, metaphase, or telophase, as well as to its state of differentiation, e.g., undifferentiated, partially differentiated, or fully differentiated. Without wanting to be limited, differentiation of a cell is usually accompanied by a decrease in the proliferative rate of a cell.

"Homology" or "identity" or "similarity" refers to sequence similarity between two peptides or between two nucleic acid molecules, with identity being a more strict comparison. Homology and identity can each be determined by comparing a position in each sequence which may be aligned for purposes of comparison. When a position in the compared sequence is occupied by the same base or amino acid, then the molecules are identical at that position. A degree of homology or similarity or

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identity between nucleic acid sequences is a function of the number of identical or matching nucleotides at positions shared by the nucleic acid sequences. A degree of identity of amino acid sequences is a function of the number of identical amino acids at positions shared by the amino acid sequences. A degree of homology or similarity of amino acid sequences is a function of the number of amino acids, i.e., structurally related, at positions shared by the amino acid sequences. An "unrelated" or "non-homologous" sequence shares less than 40% identity, though preferably less than 25% identity, with one of the sequences of the present invention.

The term "percent identical" refers to sequence identity between two amino acid sequences or between two nucleotide sequences. Identity can each be determined by comparing a position in each sequence which may be aligned for purposes of comparison. When an equivalent position in the compared sequences is occupied by the same base or amino acid, then the molecules are identical at that position; when the equivalent site occupied by the same or a similar amino acid residue (e.g., similar in steric and/or electronic nature), then the molecules can be referred to as homologous (similar) at that position. Expression as a percentage of homology, similarity, or identity refers to a function of the number of identical or similar amino acids at positions shared by the compared sequences. Various alignment algorithms and/or programs may be used, including FASTA, BLAST, or ENTREZ. FASTA and BLAST are available as a part of the GCG sequence analysis package (University of Wisconsin, Madison, Wis.), and can be used with, e.g., default settings. ENTREZ is available through the National Center for Biotechnology Information, National Library of Medicine, National Institutes of Health, Bethesda, Md. In one embodiment, the percent identity of two sequences can be determined by the GCG program with a gap weight of 1, e.g., each amino acid gap is weighted as if it were a single amino acid or nucleotide mismatch between the two sequences.

Other techniques for alignment are described in Methods in Enzymology, vol. 266: Computer Methods for Macromolecular Sequence Analysis (1996), ed. Doolittle, Academic Press, Inc., a division of Harcourt Brace & Co., San Diego, California, USA. Preferably, an alignment program that permits gaps in the sequence is utilized to align the sequences. The Smith-Waterman is one type of algorithm that permits gaps in sequence alignments. See Meth. Mol. Biol. 70: 173-187 (1997). Also, the GAP program using the Needleman and Wunsch alignment method can be utilized to

align sequences. An alternative search strategy uses MPSRCH software, which runs on a MASPAR computer. MPSRCH uses a Smith-Waterman algorithm to score sequences on a massively parallel computer. This approach improves ability to pick up distantly related matches, and is especially tolerant of small gaps and nucleotide sequence errors. Nucleic acid-encoded amino acid sequences can be used to search both protein and DNA databases.

Databases with individual sequences are described in <u>Methods in Enzymology</u>, ed. Doolittle, *supra*. Databases include Genbank, EMBL, and DNA Database of Japan (DDBJ).

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Preferred nucleic acids have a sequence at least 70%, and more preferably 80% identical and more preferably 90% and even more preferably at least 95% identical to an nucleic acid sequence of a sequence shown in one of SEQ ID NOS: 1-544. Nucleic acids at least 90%, more preferably 95%, and most preferably at least about 98-99% identical with a nucleic sequence represented in one of SEQ ID NOS: 1-544 are of course also within the scope of the invention. In preferred embodiments, the nucleic acid is mammalian.

The term "interact" as used herein is meant to include detectable interactions (e.g., biochemical interactions) between molecules, such as interaction between protein-protein, protein-nucleic acid, nucleic acid-nucleic acid, and protein-small molecule or nucleic acid-small molecule in nature.

The term "isolated" as used herein with respect to nucleic acids, such as DNA or RNA, refers to molecules separated from other DNAs, or RNAs, respectively, that are present in the natural source of the macromolecule. The term isolated as used herein also refers to a nucleic acid or peptide that is substantially free of cellular material, viral material, or culture medium when produced by recombinant DNA techniques, or chemical precursors or other chemicals when chemically synthesized. Moreover, an "isolated nucleic acid" is meant to include nucleic acid fragments which are not naturally occurring as fragments and would not be found in the natural state. The term "isolated" is also used herein to refer to polypeptides which are isolated from other cellular proteins and is meant to encompass both purified and recombinant polypeptides.

The terms "modulated" and "differentially regulated" as used herein refer to both upregulation (i.e., activation or stimulation (e.g., by agonizing or potentiating))

and downregulation (i.e., inhibition or suppression (e.g., by antagonizing, decreasing or inhibiting)).

The term "mutated gene" refers to an allelic form of a gene, which is capable of altering the phenotype of a subject having the mutated gene relative to a subject which does not have the mutated gene. If a subject must be homozygous for this mutation to have an altered phenotype, the mutation is said to be recessive. If one copy of the mutated gene is sufficient to alter the genotype of the subject, the mutation is said to be dominant. If a subject has one copy of the mutated gene and has a phenotype that is intermediate between that of a homozygous and that of a heterozygous subject (for that gene), the mutation is said to be co-dominant.

The designation "N", where it appears in the accompanying Sequence Listing, indicates that the identity of the corresponding nucleotide is unknown. "N" should therefore not necessarily be interpreted as permitting substitution with any nucleotide, e.g., A, T, C, or G, but rather as holding the place of a nucleotide whose identity has not been conclusively determined.

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The "non-human animals" of the invention include mammalians such as rodents, non-human primates, sheep, dog, cow, chickens, amphibians, reptiles, etc. Preferred non-human animals are selected from the rodent family including rat and mouse, most preferably mouse, though transgenic amphibians, such as members of the *Xenopus* genus, and transgenic chickens can also provide important tools for understanding and identifying agents which can affect, for example, embryogenesis and tissue formation. The term "chimeric animal" is used herein to refer to animals in which the recombinant gene is found, or in which the recombinant gene is expressed in some but not all cells of the animal. The term "tissue-specific chimeric animal" indicates that one of the recombinant genes is present and/or expressed or disrupted in some tissues but not others.

As used herein, the term "nucleic acid" refers to polynucleotides such as deoxyribonucleic acid (DNA), and, where appropriate, ribonucleic acid (RNA). The term should also be understood to include, as equivalents, analogs of either RNA or DNA made from nucleotide analogs, and, as applicable to the embodiment being described, single (sense or antisense) and double-stranded polynucleotides. ESTs, chromosomes, cDNAs, mRNAs, and rRNAs are representative examples of molecules that may be referred to as nucleic acids.

The term "nucleotide sequence complementary to the nucleotide sequence of SEQ ID NO. x" refers to the nucleotide sequence of the complementary strand of a nucleic acid strand having SEQ ID NO. x. The term "complementary strand" is used herein interchangeably with the term "complement". The complement of a nucleic acid strand can be the complement of a coding strand or the complement of a non-coding strand.

The term "polymorphism" refers to the coexistence of more than one form of a gene or portion (e.g., allelic variant) thereof. A portion of a gene of which there are at least two different forms, i.e., two different nucleotide sequences, is referred to as a "polymorphic region of a gene". A polymorphic region can be a single nucleotide, the identity of which differs in different alleles. A polymorphic region can also be several nucleotides long.

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A "polymorphic gene" refers to a gene having at least one polymorphic region.

As used herein, the term "promoter" means a DNA sequence that regulates expression of a selected DNA sequence operably linked to the promoter, and which effects expression of the selected DNA sequence in cells. The term encompasses "tissue specific" promoters, i.e., promoters which effect expression of the selected DNA sequence only in specific cells (e.g., cells of a specific tissue). The term also covers so-called "leaky" promoters, which regulate expression of a selected DNA primarily in one tissue, but cause expression in other tissues as well. The term also encompasses non-tissue specific promoters and promoters that constitutively expressed or that are inducible (i.e., expression levels can be controlled).

The terms "protein", "polypeptide", and "peptide" are used interchangeably herein when referring to a gene product.

The term "recombinant protein" refers to a polypeptide of the present invention which is produced by recombinant DNA techniques, wherein generally, DNA encoding a polypeptide is inserted into a suitable expression vector which is in turn used to transform a host cell to produce the heterologous protein. Moreover, the phrase "derived from", with respect to a recombinant gene, is meant to include within the meaning of "recombinant protein" those proteins having an amino acid sequence of a native polypeptide, or an amino acid sequence similar thereto which is generated by mutations including substitutions and deletions (including truncation) of a naturally occurring form of the polypeptide.

"Small molecule" as used herein, is meant to refer to a composition, which has a molecular weight of less than about 5 kD and most preferably less than about 4 kD. Small molecules can be nucleic acids, peptides, polypeptides, peptidomimetics, carbohydrates, lipids or other organic (carbon-containing) or inorganic molecules. Many pharmaceutical companies have extensive libraries of chemical and/or biological mixtures, often fungal, bacterial, or algal extracts, which can be screened with any of the assays of the invention to identify compounds that modulate a bioactivity.

As used herein, the term "specifically hybridizes" or "specifically detects" refers to the ability of a nucleic acid molecule of the invention to hybridize to at least a portion of, for example, approximately 6, 12, 15, 20, 30, 50, 100, 150, 200, 300, 350, 400, 500, 750, or 1000 contiguous nucleotides of a nucleic acid designated in any one of SEQ ID Nos: 1-544, or a sequence complementary thereto, or naturally occurring mutants thereof, such that it has less than 15%, preferably less than 10%, and more preferably less than 5% background hybridization to a cellular nucleic acid (e.g., mRNA or genomic DNA) encoding a different protein. In preferred embodiments, the oligonucleotide probe detects only a specific nucleic acid, e.g., it does not substantially hybridize to similar or related nucleic acids, or complements thereof.

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"Transcriptional regulatory sequence" is a generic term used throughout the specification to refer to DNA sequences, such as initiation signals, enhancers, and promoters, which induce or control transcription of protein coding sequences with which they are operably linked. In preferred embodiments, transcription of one of the genes is under the control of a promoter sequence (or other transcriptional regulatory sequence) which controls the expression of the recombinant gene in a cell-type in which expression is intended. It will also be understood that the recombinant gene can be under the control of transcriptional regulatory sequences which are the same or which are different from those sequences which control transcription of the naturally-occurring forms of the polypeptide.

As used herein, the term "transfection" means the introduction of a nucleic acid, e.g., via an expression vector, into a recipient cell by nucleic acid-mediated gene transfer. "Transformation", as used herein, refers to a process in which a cell's genotype is changed as a result of the cellular uptake of exogenous DNA or RNA,

and, for example, the transformed cell expresses a recombinant form of a polypeptide or, in the case of anti-sense expression from the transferred gene, the expression of the target gene is disrupted.

As used herein, the term "transgene" means a nucleic acid sequence (or an antisense transcript thereto) which has been introduced into a cell. A transgene could be partly or entirely heterologous, i.e., foreign, to the transgenic animal or cell into which it is introduced, or, is homologous to an endogenous gene of the transgenic animal or cell into which it is introduced, but which is designed to be inserted, or is inserted, into the animal's genome in such a way as to alter the genome of the cell into which it is inserted (e.g., it is inserted at a location which differs from that of the natural gene or its insertion results in a knockout). A transgene can also be present in a cell in the form of an episome. A transgene can include one or more transcriptional regulatory sequences and any other nucleic acid, such as introns, that may be necessary for optimal expression of a selected nucleic acid.

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A "transgenic animal" refers to any animal, preferably a non-human mammal, bird or an amphibian, in which one or more of the cells of the animal contain heterologous nucleic acid introduced by way of human intervention, such as by transgenic techniques well known in the art. The nucleic acid is introduced into the cell, directly or indirectly by introduction into a precursor of the cell, by way of deliberate genetic manipulation, such as by microinjection or by infection with a recombinant virus. The term genetic manipulation does not include classical crossbreeding, or in vitro fertilization, but rather is directed to the introduction of a recombinant DNA molecule. This molecule may be integrated within a chromosome, or it may be extra-chromosomally replicating DNA. In the typical transgenic animals described herein, the transgene causes cells to express a recombinant form of one of the subject polypeptide, e.g. either agonistic or antagonistic forms. However, transgenic animals in which the recombinant gene is silent are also contemplated, as for example, the FLP or CRE recombinase dependent constructs described below. Moreover, "transgenic animal" also includes those recombinant animals in which gene disruption of one or more genes is caused by human intervention, including both recombination and antisense techniques.

The term "treating" as used herein is intended to encompass curing as well as ameliorating at least one symptom of the condition or disease.

The term "vector" refers to a nucleic acid molecule capable of transporting another nucleic acid to which it has been linked. One type of preferred vector is an episome, i.e., a nucleic acid capable of extra-chromosomal replication. Preferred vectors are those capable of autonomous replication and/or expression of nucleic acids to which they are linked. Vectors capable of directing the expression of genes to which they are operatively linked are referred to herein as "expression vectors". In general, expression vectors of utility in recombinant DNA techniques are often in the form of "plasmids" which refer generally to circular double stranded DNA loops which, in their vector form are not bound to the chromosome. In the present specification, "plasmid" and "vector" are used interchangeably as the plasmid is the most commonly used form of vector. However, the invention is intended to include such other forms of expression vectors which serve equivalent functions and which become known in the art subsequently hereto.

The term "wild-type allele" refers to an allele of a gene which, when present in two copies in a subject results in a wild-type phenotype. There can be several different wild-type alleles of a specific gene, since certain nucleotide changes in a gene may not affect the phenotype of a subject having two copies of the gene with the nucleotide changes.

#### 20 III. Nucleic Acids of the Present Invention

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As described below, one aspect of the invention pertains to isolated nucleic acids, variants, and/or equivalents of such nucleic acids.

Nucleic acids of the present invention have been identified as differentially expressed in tumor cells, e.g., colon cancer-derived cell lines (relative to the expression levels in normal tissue, e.g., normal colon tissue and/or normal non-colon tissue), such as SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto. In certain embodiments, the subject nucleic acids are differentially expressed by at least a factor of two, preferably at least a factor of five, even more preferably at least a factor of twenty, still more preferably at least a factor of fifty. Preferred nucleic acids include sequences identified as differentially expressed both in colon cancer cell tissue and colon cancer cell lines. In preferred embodiments, nucleic acids of the present invention are upregulated in tumor cells, especially colon cancer tissue and/or colon

cancer-derived cell lines. In another embodiment, nucleic acids of the present invention are downregulated in tumor cells, especially colon cancer tissue and/or colon cancer-derived cell lines.

Table 1 indicates those sequences which are over- or underexpressed in a colon cancer-derived cell line relative to normal tissue, and further designates those sequences which are also differentially regulated in colon cancer tissue. The designation O indicates that the corresponding sequence was overexpressed, M indicates possible overexpression, N indicates no differential expression, and U indicates underexpression.

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Genes which are upregulated, such as oncogenes, or downregulated, such as tumor suppressors, in aberrantly proliferating cells may be targets for diagnostic or therapeutic techniques. For example, upregulation of the *cdc2* gene induces mitosis. Overexpression of the *myt1* gene, a mitotic deactivator, negatively regulates the activity of *cdc2*. Aberrant proliferation may thus be induced either by upregulating *cdc2* or by downregulating *myt1*. Similarly, downregulation of tumor suppressors such as *p53* and *Rb* have been implicated in tumorigenesis.

Particularly preferred polypeptides are those that are encoded by nucleic acid sequences at least about 70%, 75%, 80%, 90%, 95%, 97%, or 98% similar to a nucleic acid sequence of SEQ ID Nos. 1-544. Preferably, the nucleic acid includes all or a portion (e.g., at least about 12, at least about 15, at least about 25, or at least about 40 nucleotides) of the nucleotide sequence corresponding to the nucleic acid of SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35, or a sequence complementary thereto.

Still other preferred nucleic acids of the present invention encode a polypeptide comprising at least a portion of a polypeptide encoded by one of SEQ ID Nos. 1-544. For example, preferred nucleic acid molecules for use as probes/primers or antisense molecules (i.e., noncoding nucleic acid molecules) can comprise at least about 12, 20, 30, 50, 60, 70, 80, 90, or 100 base pairs in length up to the length of the complete gene. Coding nucleic acid molecules can comprise, for example, from about 50, 60, 70, 80, 90, or 100 base pairs up to the length of the complete gene.

Another aspect of the invention provides a nucleic acid which hybridizes under low, medium, or high stringency conditions to a nucleic acid sequence represented by one of SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35, or a sequence complementary thereto. Appropriate stringency conditions which promote

DNA hybridization, for example, 6.0 x sodium chloride/sodium citrate (SSC) at about 45 °C, followed by a wash of 2.0 x SSC at 50 °C, are known to those skilled in the art or can be found in Current Protocols in Molecular Biology, John Wiley & Sons, N.Y. (1989), 6.3.1-12.3.6. For example, the salt concentration in the wash step can be selected from a low stringency of about 2.0 x SSC at 50 °C to a high stringency of about 0.2 x SSC at 50 °C. In addition, the temperature in the wash step can be increased from low stringency conditions at room temperature, about 22 °C, to high stringency conditions at about 65 °C. Both temperature and salt may be varied, or temperature or salt concentration may be held constant while the other variable is changed. In a preferred embodiment, a nucleic acid of the present invention will bind to one of SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, under moderately stringent conditions, for example at about 2.0 x SSC and about 40 °C. In a particularly preferred embodiment, a nucleic acid of the present invention will bind to one of SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, under high stringency conditions.

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In one embodiment, the invention provides nucleic acids which hybridize under low stringency conditions of  $6 \times SSC$  at room temperature followed by a wash at  $2 \times SSC$  at room temperature.

In another embodiment, the invention provides nucleic acids which hybridize under high stringency conditions of 2 x SSC at 65 °C followed by a wash at 0.2 x SSC at 65 °C.

Nucleic acids having a sequence that differs from the nucleotide sequences shown in one of SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, due to degeneracy in the genetic code, are also within the scope of the invention. Such nucleic acids encode functionally equivalent peptides (i.e., a peptide having equivalent or similar biological activity) but differ in sequence from the sequence shown in the sequence listing due to degeneracy in the genetic code. For example, a number of amino acids are designated by more than one triplet. Codons that specify the same amino acid, or synonyms (for example, CAU and CAC each encode histidine) may result in "silent" mutations which do not affect the amino acid sequence of a polypeptide. However, it is expected that DNA sequence polymorphisms that do lead to changes in the amino acid sequences of the subject polypeptides will exist among mammals. One skilled in the art will appreciate that

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these variations in one or more nucleotides (e.g., up to about 3-5% of the nucleotides) of the nucleic acids encoding polypeptides having an activity of a polypeptide may exist among individuals of a given species due to natural allelic variation.

Also within the scope of the invention are nucleic acids encoding splicing variants of proteins encoded by a nucleic acid of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, or natural homologs of such proteins. Such homologs can be cloned by hybridization or PCR, as further described herein.

The polynucleotide sequence may also encode for a leader sequence, e.g., the natural leader sequence or a heterologous leader sequence, for a subject polypeptide. For example, the desired DNA sequence may be fused in the same reading frame to a DNA sequence which aids in expression and secretion of the polypeptide from the host cell, for example, a leader sequence which functions as a secretory sequence for controlling transport of the polypeptide from the cell. The protein having a leader sequence is a preprotein and may have the leader sequence cleaved by the host cell to form the mature form of the protein.

The polynucleotide of the present invention may also be fused in frame to a marker sequence, also referred to herein as "Tag sequence" encoding a "Tag peptide", which allows for marking and/or purification of the polypeptide of the present invention. In a preferred embodiment, the marker sequence is a hexahistidine tag, e.g., supplied by a PQE-9 vector. Numerous other Tag peptides are available commercially. Other frequently used Tags include myc-epitopes (e.g., see Ellison et al. (1991) *J Biol Chem 266*:21150-21157) which includes a 10-residue sequence from c-myc, the pFLAG system (International Biotechnologies, Inc.), the pEZZ-protein A system (Pharmacia, NJ), and a 16 amino acid portion of the *Haemophilus influenza* hemagglutinin protein. Furthermore, any polypeptide can be used as a Tag so long as a reagent, e.g., an antibody interacting specifically with the Tag polypeptide is available or can be prepared or identified.

As indicated by the examples set out below, nucleic acids can be obtained from mRNA present in any of a number of eukaryotic cells, e.g., and are preferably obtained from metazoan cells, more preferably from vertebrate cells, and even more preferably from mammalian cells. It should also be possible to obtain nucleic acids of the present invention from genomic DNA from both adults and embryos. For

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example, a gene can be cloned from either a cDNA or a genomic library in accordance with protocols generally known to persons skilled in the art. cDNA can be obtained by isolating total mRNA from a cell, e.g., a vertebrate cell, a mammalian cell, or a human cell, including embryonic cells. Double stranded cDNAs can then be prepared from the total mRNA, and subsequently inserted into a suitable plasmid or bacteriophage vector using any one of a number of known techniques. The gene can also be cloned using established polymerase chain reaction techniques in accordance with the nucleotide sequence information provided by the invention.

In certain embodiments, a nucleic acid, probe, vector, or other construct of the present invention includes at least about five, at least about ten, or at least about twenty nucleic acids from a region designated as novel in Table 2. In certain other embodiments, a nucleic acid of the present invention includes at least about five, at least about ten, or at least about twenty nucleic acids which are not included in the clones whose accession numbers are listed in Table 2.

The invention includes within its scope a polynucleotide having the nucleotide sequence of nucleic acid obtained from this biological material, wherein the nucleic acid hybridizes under stringent conditions (at least about 4 x SSC at 65 °C, or at least about 4 x SSC at 42 °C; see, for example, U.S. Patent No. 5,707,829, incorporated herein by reference) with at least 15 contiguous nucleotides of at least one of SEQ ID Nos. 1-544. By this is intended that when at least 15 contiguous nucleotides of one of SEQ ID Nos. 1-544 is used as a probe, the probe will preferentially hybridize with a gene or mRNA (of the biological material) comprising the complementary sequence, allowing the identification and retrieval of the nucleic acids of the biological material that uniquely hybridize to the selected probe. Probes from more than one of SEQ ID Nos. 1-544 will hybridize with the same gene or mRNA if the cDNA from which they were derived corresponds to one mRNA. Probes of more than 15 nucleotides can be used, but 15 nucleotides represents enough sequence for unique identification.

Because the present nucleic acids represent partial mRNA transcripts, two or more nucleic acids of the invention may represent different regions of the same mRNA transcript and the same gene. Thus, if two or more of SEQ ID Nos. 1-544 are identified as belonging to the same clone, then either sequence can be used to obtain the full-length mRNA or gene.

Nucleic acid-related polynucleotides can also be isolated from cDNA libraries. These libraries are preferably prepared from mRNA of human colon cells, more preferably, human colon cancer specific tissue, designated as the DE clones in the appended Tables. In another embodiment the nucleic acids are isolated from libraries prepared from normal colon specific tissue, designated herein as PA clones in the appended Tables. In yet another embodiment, this invention discloses nucleic acid sequences that can be isolated from both libraries prepared from a human colon adenocarcinoma cell line, SW480, as well as from libraries prepared from either normal colon specific tissue or from colon cancer specific tissue. These sequences are listed in Table 3. Alignment of SEQ ID Nos. 1-544, as described above, can indicated that a cell line or tissue source of a related protein or polynucleotide can also be used as a source of the nucleic acid-related cDNA.

Techniques for producing and probing nucleic acid sequence libraries are described, for example, in Sambrook et al., "Molecular Cloning: A Laboratory Manual" (New York, Cold Spring Harbor Laboratory, 1989). The cDNA can be prepared by using primers based on a sequence from SEQ ID Nos. 1-544. In one embodiment, the cDNA library can be made from only poly-adenylated mRNA. Thus, poly-T primers can be used to prepare cDNA from the mRNA. Alignment of SEQ ID Nos. 1-544 can result in identification of a related polypeptide or polynucleotide. Some of the polynucleotides disclosed herein contains repetitive regions that were subject to masking during the search procedures. The information about the repetitive regions is discussed below.

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Constructs of polynucleotides having sequences of SEQ ID Nos. 1-544 can be generated synthetically. Alternatively, single-step assembly of a gene and entire plasmid from large numbers of oligodeoxyribonucleotides is described by Stemmer et al., *Gene (Amsterdam)* (1995) 164(1):49-53. In this method, assembly PCR (the synthesis of long DNA sequences from large numbers of oligodeoxyribonucleotides (oligos)) is described. The method is derived from DNA shuffling (Stemmer, *Nature* (1994) 370:389-391), and does not rely on DNA ligase, but instead relies on DNA polymerase to build increasingly longer DNA fragments during the assembly process. For example, a 1.1-kb fragment containing the TEM-1 beta-lactamase-encoding gene (bla) can be assembled in a single reaction from a total of 56 oligos, each 40 nucleotides (nt) in length. The synthetic gene can be PCR amplified and cloned in a

vector containing the tetracycline-resistance gene (Tc-R) as the sole selectable marker. Without relying on ampicillin (Ap) selection, 76% of the Tc-R colonies were Ap-R, making this approach a general method for the rapid and cost-effective synthesis of any gene.

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# IV. <u>Identification of Functional and Structural Motifs of Novel-Genes Using Art-Recognized Methods</u>

Translations of the nucleotide sequence of the nucleic acids, cDNAs, or full genes can be aligned with individual known sequences. Similarity with individual sequences can be used to determine the activity of the polypeptides encoded by the polynucleotides of the invention. For example, sequences that show similarity with a chemokine sequence may exhibit chemokine activities. Also, sequences exhibiting similarity with more than one individual sequence may exhibit activities that are characteristic of either or both individual sequences.

The full length sequences and fragments of the polynucleotide sequences of the nearest neighbors can be used as probes and primers to identify and isolate the full length sequence of the nucleic acid. The nearest neighbors can indicate a tissue or cell type to be used to construct a library for the full-length sequences of the nucleic acid.

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Typically, the nucleic acids are translated in all six frames to determine the best alignment with the individual sequences. The sequences disclosed herein in the Sequence Listing are in a 5' to 3' orientation and translation in three frames can be sufficient (with a few specific exceptions as described in the Examples). These amino acid sequences are referred to, generally, as query sequences, which will be aligned with the individual sequences.

Nucleic acid sequences can be compared with known genes by any of the methods disclosed above. Results of individual and query sequence alignments can be divided into three categories: high similarity, weak similarity, and no similarity. Individual alignment results ranging from high similarity to weak similarity provide a basis for determining polypeptide activity and/or structure.

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Parameters for categorizing individual results include: percentage of the alignment region length where the strongest alignment is found, percent sequence identity, and p value.

The percentage of the alignment region length is calculated by counting the number of residues of the individual sequence found in the region of strongest alignment. This number is divided by the total residue length of the query sequence to find a percentage. An example is shown below:

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Query sequence: ASNPERTMIPVTRVGLIRYM

Individual sequence: YMMTEYLAIPV.RVGLPRYM

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The region of alignment begins at amino acid 9 and ends at amino acid 19. The total length of the query sequence is 20 amino acids. The percent of the alignment region length is 11/20 or 55%.

Percent sequence identity is calculated by counting the number of amino acid matches between the query and individual sequence and dividing total number of matches by the number of residues of the individual sequence found in the region of strongest alignment. For the example above, the percent identity would be 10 matches divided by 11 amino acids, or approximately 90.9%.

P value is the probability that the alignment was produced by chance. For a single alignment, the p value can be calculated according to Karlin et al., Proc. Natl. Acad. Sci. 87: 2264 (1990) and Karlin et al., Proc. Natl. Acad. Sci. 90: (1993). The p value of multiple alignments using the same query sequence can be calculated using an heuristic approach described in Altschul et al., Nat. Genet. 6: 119 (1994).

Alignment programs such as BLAST program can calculate the p value.

The boundaries of the region where the sequences align can be determined according to Doolittle, Methods in Enzymology, *supra*; BLAST or FASTA programs; or by determining the area where the sequence identity is highest.

Another factor to consider for determining identity or similarity is the location of the similarity or identity. Strong local alignment can indicate similarity even if the length of alignment is short. Sequence identity scattered throughout the length of the query sequence also can indicate a similarity between the query and profile sequences.

#### High Similarity

For the alignment results to be considered high similarity, the percent of the alignment region length, typically, is at least about 55% of total length query sequence; more typically, at least about 58%; even more typically; at least about 60% of the total residue length of the query sequence. Usually, percent length of the alignment region can be as much as about 62%; more usually, as much as about 64%; even more usually, as much as about 66%.

Further, for high similarity, the region of alignment, typically, exhibits at least about 75% of sequence identity; more typically, at least about 78%; even more typically; at least about 80% sequence identity. Usually, percent sequence identity can be as much as about 82%; more usually, as much as about 84%; even more usually, as much as about 86%.

The p value is used in conjunction with these methods. If high similarity is found, the query sequence is considered to have high similarity with a profile sequence when the p value is less than or equal to about  $10^{-2}$ ; more usually; less than or equal to about  $10^{-3}$ ; even more usually; less than or equal to about  $10^{-4}$ . More typically, the p value is no more than about  $10^{-5}$ ; more typically; no more than or equal to about  $10^{-15}$  for the query sequence to be considered high similarity.

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#### Weak Similarity

For the alignment results to be considered weak similarity, there is no minimum percent length of the alignment region nor minimum length of alignment. A better showing of weak similarity is considered when the region of alignment is, typically, at least about 15 amino acid residues in length; more typically, at least about 20; even more typically; at least about 25 amino acid residues in length. Usually, length of the alignment region can be as much as about 30 amino acid residues; more usually, as much as about 40; even more usually, as much as about 60 amino acid residues.

Further, for weak similarity, the region of alignment, typically, exhibits at least about 35% of sequence identity; more typically, at least about 40%; even more typically; at least about 45% sequence identity. Usually, percent sequence identity

can be as much as about 50%; more usually, as much as about 55%; even more usually, as much as about 60%.

If low similarity is found, the query sequence is considered to have weak similarity with a profile sequence when the p value is usually less than or equal to about  $10^{-2}$ ; more usually; less than or equal to about  $10^{-3}$ ; even more usually; less than or equal to about  $10^{-4}$ . More typically, the p value is no more than about  $10^{-5}$ ; more usually; no more than or equal to about  $10^{-10}$ ; even more usually; no more than or equal to about  $10^{-15}$  for the query sequence to be considered weak similarity.

#### 10 Similarity Determined by Sequence Identity

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Sequence identity alone can be used to determine similarity of a query sequence to an individual sequence and can indicate the activity of the sequence. Such an alignment, preferably, permits gaps to align sequences. Typically, the query sequence is related to the profile sequence if the sequence identity over the entire query sequence is at least about 15%; more typically, at least about 20%; even more typically, at least about 25%; even more typically, at least about 50%. Sequence identity alone as a measure of similarity is most useful when the query sequence is usually, at least 80 residues in length; more usually, 90 residues; even more usually, at least 95 amino acid residues in length. More typically, similarity can be concluded based on sequence identity alone when the query sequence is preferably 100 residues in length; more preferably, 120 residues in length; even more preferably, 150 amino acid residues in length.

## Determining Activity from Alignments with Profile and Multiple Aligned Sequences

Translations of the nucleic acids can be aligned with amino acid profiles that define either protein families or common motifs. Also, translations of the nucleic acids can be aligned to multiple sequence alignments (MSA) comprising the polypeptide sequences of members of protein families or motifs. Similarity or identity with profile sequences or MSAs can be used to determine the activity of the polypeptides encoded by nucleic acids or corresponding cDNA or genes. For example, sequences that show an identity or similarity with a chemokine profile or MSA can exhibit chemokine activities.

Profiles can designed manually by (1) creating a MSA, which is an alignment of the amino acid sequence of members that belong to the family and (2) constructing

a statistical representation of the alignment. Such methods are described, for example, in Birney et al., Nucl. Acid Res. 24(14): 2730-2739 (1996).

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MSAs of some protein families and motifs are publicly available. For example, these include MSAs of 547 different families and motifs. These MSAs are described also in Sonnhammer et al., <u>Proteins 28</u>: 405-420 (1997). Other sources are also available in the world wide web. A brief description of these MSAs is reported in Pascarella et al., <u>Prot. Eng. 9(3)</u>: 249-251 (1996).

Techniques for building profiles from MSAs are described in Sonnhammer et al., supra; Birney et al., supra; and Methods in Enzymology, vol. 266: "Computer Methods for Macromolecular Sequence Analysis," 1996, ed. Doolittle, Academic Press, Inc., a division of Harcourt Brace & Co., San Diego, California, USA.

Similarity between a query sequence and a protein family or motif can be determined by (a) comparing the query sequence against the profile and/or (b) aligning the query sequence with the members of the family or motif.

Typically, a program such as Searchwise can be used to compare the query sequence to the statistical representation of the multiple alignment, also known as a profile. The program is described in Birney et al., supra. Other techniques to compare the sequence and profile are described in Sonnhammer et al., supra and Doolittle, supra.

Next, methods described by Feng et al., <u>J. Mol. Evol. 25</u>: 351-360 (1987) and Higgins et al., <u>CABIOS</u> 5: 151-153 (1989) can be used align the query sequence with the members of a family or motif, also known as a MSA. Computer programs, such as PILEUP, can be used. See Feng et al., infra.

The following factors are used to determine if a similarity between a query sequence and a profile or MSA exists: (1) number of conserved residues found in the query sequence, (2) percentage of conserved residues found in the query sequence, (3) number of frameshifts, and (4) spacing between conserved residues.

Some alignment programs that both translate and align sequences can make any number of frameshifts when translating the nucleotide sequence to produce the best alignment. The fewer frameshifts needed to produce an alignment, the stronger the similarity or identity between the query and profile or MSAs. For example, a weak similarity resulting from no frameshifts can be a better indication of activity or structure of a query sequence, than a strong similarity resulting from two frameshifts.

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Preferably, three or fewer frameshifts are found in an alignment; more preferably two or fewer frameshifts; even more preferably, one or fewer frameshifts; even more preferably, no frameshifts are found in an alignment of query and profile or MSAs.

Conserved residues are those amino acids that are found at a particular position in all or some of the family or motif members. For example, most known chemokines contain four conserved cysteines. Alternatively, a position is considered conserved if only a certain class of amino acids is found in a particular position in all or some of the family members. For example, the N-terminal position may contain a positively charged amino acid, such as lysine, arginine, or histidine.

Typically, a residue of a polypeptide is conserved when a class of amino acids or a single amino acid is found at a particular position in at least about 40% of all class members; more typically, at least about 50%; even more typically, at least about 60% of the members. Usually, a residue is conserved when a class or single amino acid is found in at least about 70% of the members of a family or motif; more usually, at least about 80%; even more usually, at least about 90%; even more usually, at least about 95%.

A residue is considered conserved when three unrelated amino acids are found at a particular position in the some or all of the members; more usually, two unrelated amino acids. These residues are conserved when the unrelated amino acids are found at particular positions in at least about 40% of all class member; more typically, at least about 50%; even more typically, at least about 60% of the members. Usually, a residue is conserved when a class or single amino acid is found in at least about 70% of the members of a family or motif; more usually, at least about 80%; even more usually, at least about 95%.

A query sequence has similarity to a profile or MSA when the query sequence comprises at least about 25% of the conserved residues of the profile or MSA; more usually, at least about 30%; even more usually; at least about 40%. Typically, the query sequence has a stronger similarity to a profile sequence or MSA when the query sequence comprises at least about 45% of the conserved residues of the profile or MSA; more typically, at least about 50%; even more typically; at least about 55%.

#### V. Probes and Primers

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The nucleotide sequences determined from the cloning of genes from tumor cells, especially colon cancer cell lines and tissues will further allow for the generation of probes and primers designed for identifying and/or cloning homologs in other cell types, e.g., from other tissues, as well as homologs from other mammalian organisms. Nucleotide sequences useful as probes/primers may include all or a portion of the sequences listed in SEQ ID Nos. 1-544 or sequences complementary thereto or sequences which hybridize under stringent conditions to all or a portion of SEQ ID Nos. 1-544. For instance, the present invention also provides a probe/primer comprising a substantially purified oligonucleotide, which oligonucleotide comprising a nucleotide sequence that hybridizes under stringent conditions to at least approximately 12, preferably 25, more preferably 40, 50, or 75 consecutive nucleotides up to the full length of the sense or anti-sense sequence selected from the group consisting of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, or naturally occurring mutants thereof. For instance, primers based on a nucleic acid represented in SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, can be used in PCR reactions to clone homologs of that sequence.

In yet another embodiment, the invention provides probes/primers comprising a nucleotide sequence that hybridizes under moderately stringent conditions to at least approximately 12, 16, 25, 40, 50 or 75 consecutive nucleotides up to the full length of the sense or antisense sequence selected from the group consisting of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or naturally occurring mutants thereof.

In particular, these probes are useful because they provide a method for detecting mutations in wild-type genes of the present invention. Nucleic acid probes which are complementary to a wild-type gene of the present invention and can form mismatches with mutant genes are provided, allowing for detection by enzymatic or chemical cleavage or by shifts in electrophoretic mobility.

Likewise, probes based on the subject sequences can be used to detect transcripts or genomic sequences encoding the same or homologous proteins, for use, for example, in prognostic or diagnostic assays. In preferred embodiments, the probe

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further comprises a label group attached thereto and able to be detected, e.g., the label group is selected from radioisotopes, fluorescent compounds, chemiluminescent compounds, enzymes, and enzyme co-factors.

Full-length cDNA molecules comprising the disclosed nucleic acids are obtained as follows. A subject nucleic acid or a portion thereof comprising at least about 12, 15, 18, or 20 nucleotides up to the full length of a sequence represented in SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, may be used as a hybridization probe to detect hybridizing members of a cDNA library using probe design methods, cloning methods, and clone selection techniques as described in U.S. Patent No. 5,654,173, "Secreted Proteins and Polynucleotides Encoding Them," incorporated herein by reference. Libraries of cDNA may be made from selected tissues, such as normal or tumor tissue, or from tissues of a mammal treated with, for example, a pharmaceutical agent. Preferably, the tissue is the same as that used to generate the nucleic acids, as both the nucleic acid and the cDNA represent expressed genes. Most preferably, the cDNA library is made from the biological material described herein in the Examples. Alternatively, many cDNA libraries are available commercially. (Sambrook et al., Molecular Cloning: A Laboratory Manual, 2nd Ed. (Cold Spring Harbor Press, Cold Spring Harbor, NY 1989). The choice of cell type for library construction may be made after the identity of the protein encoded by the nucleic acid-related gene is known. This will indicate which tissue and cell types are likely to express the related gene, thereby containing the mRNA for generating the cDNA.

Members of the library that are larger than the nucleic acid, and preferably that contain the whole sequence of the native message, may be obtained. To confirm that the entire cDNA has been obtained, RNA protection experiments may be performed as follows. Hybridization of a full-length cDNA to an mRNA may protect the RNA from RNase degradation. If the cDNA is not full length, then the portions of the mRNA that are not hybridized may be subject to RNase degradation. This may be assayed, as is known in the art, by changes in electrophoretic mobility on polyacrylamide gels, or by detection of released monoribonucleotides. Sambrook <u>et al.</u>, Molecular Cloning: A Laboratory Manual, 2nd Ed. (Cold Spring Harbor Press, Cold Spring Harbor, NY 1989). In order to obtain additional sequences 5' to the end

of a partial cDNA, 5' RACE (PCR Protocols: A Guide to Methods and Applications (Academic Press, Inc. 1990)) may be performed.

Genomic DNA may be isolated using nucleic acids in a manner similar to the isolation of full-length cDNAs. Briefly, the nucleic acids, or portions thereof, may be used as probes to libraries of genomic DNA. Preferably, the library is obtained from the cell type that was used to generate the nucleic acids. Most preferably, the genomic DNA is obtained from the biological material described herein in the Example. Such libraries may be in vectors suitable for carrying large segments of a genome, such as P1 or YAC, as described in detail in Sambrook <u>et al.</u>, 9.4-9.30. In addition, genomic sequences can be isolated from human BAC libraries, which are commercially available from Research Genetics, Inc., Huntville, Alabama, USA, for example. In order to obtain additional 5' or 3' sequences, chromosome walking may be performed, as described in Sambrook <u>et al.</u>, such that adjacent and overlapping fragments of genomic DNA are isolated. These may be mapped and pieced together, as is known in the art, using restriction digestion enzymes and DNA ligase.

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Using the nucleic acids of the invention, corresponding full length genes can be isolated using both classical and PCR methods to construct and probe cDNA libraries. Using either method, Northern blots, preferably, may be performed on a number of cell types to determine which cell lines express the gene of interest at the highest rate.

Classical methods of constructing cDNA libraries are taught in Sambrook et al., supra. With these methods, cDNA can be produced from mRNA and inserted into viral or expression vectors. Typically, libraries of mRNA comprising poly(A) tails can be produced with poly(T) primers. Similarly, cDNA libraries can be produced using the instant sequences as primers.

PCR methods may be used to amplify the members of a cDNA library that comprise the desired insert. In this case, the desired insert may contain sequence from the full length cDNA that corresponds to the instant nucleic acids. Such PCR methods include gene trapping and RACE methods.

Gene trapping may entail inserting a member of a cDNA library into a vector. The vector then may be denatured to produce single stranded molecules. Next, a substrate-bound probe, such a biotinylated oligo, may be used to trap cDNA inserts of interest. Biotinylated probes can be linked to an avidin-bound solid substrate. PCR

methods can be used to amplify the trapped cDNA. To trap sequences corresponding to the full length genes, the labeled probe sequence may be based on the nucleic acids of the invention, e.g., SEQ ID Nos. 1-168, preferably SEQ ID Nos. 1-35, or a sequence complementary thereto. Random primers or primers specific to the library vector can be used to amplify the trapped cDNA. Such gene trapping techniques are described in Gruber et al., PCT WO 95/04745 and Gruber et al., U.S. Pat. No. 5,500,356. Kits are commercially available to perform gene trapping experiments from, for example, Life Technologies, Gaithersburg, Maryland, USA.

"Rapid amplification of cDNA ends," or RACE, is a PCR method of amplifying cDNAs from a number of different RNAs. The cDNAs may be ligated to an oligonucleotide linker and amplified by PCR using two primers. One primer may be based on sequence from the instant nucleic acids, for which full length sequence is desired, and a second primer may comprise a sequence that hybridizes to the oligonucleotide linker to amplify the cDNA. A description of this method is reported in PCT Pub. No. WO 97/19110.

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In preferred embodiments of RACE, a common primer may be designed to anneal to an arbitrary adaptor sequence ligated to cDNA ends (Apte and Siebert, Biotechniques 15:890-893, 1993; Edwards et al., Nuc. Acids Res. 19:5227-5232, 1991). When a single gene-specific RACE primer is paired with the common primer, preferential amplification of sequences between the single gene specific primer and the common primer occurs. Commercial cDNA pools modified for use in RACE are available.

Another PCR-based method generates full-length cDNA library with anchored ends without specific knowledge of the cDNA sequence. The method uses lock-docking primers (I-VI), where one primer, poly TV (I-III) locks over the polyA tail of eukaryotic mRNA producing first strand synthesis and a second primer, polyGH (IV-VI) locks onto the polyC tail added by terminal deoxynucleotidyl transferase (TdT). This method is described in PCT Pub. No. WO 96/40998.

The promoter region of a gene generally is located 5' to the initiation site for RNA polymerase II. Hundreds of promoter regions contain the "TATA" box, a sequence such as TATTA or TATAA, which is sensitive to mutations. The promoter region can be obtained by performing 5' RACE using a primer from the coding region

of the gene. Alternatively, the cDNA can be used as a probe for the genomic sequence, and the region 5' to the coding region is identified by "walking up."

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If the gene is highly expressed or differentially expressed, the promoter from the gene may be of use in a regulatory construct for a heterologous gene.

Once the full-length cDNA or gene is obtained, DNA encoding variants can be prepared by site-directed mutagenesis, described in detail in Sambrook <u>et al.</u>, 15.3-15.63. The choice of codon or nucleotide to be replaced can be based on the disclosure herein on optional changes in amino acids to achieve altered protein structure and/or function.

As an alternative method to obtaining DNA or RNA from a biological material, nucleic acid comprising nucleotides having the sequence of one or more nucleic acids of the invention can be synthesized. Thus, the invention encompasses nucleic acid molecules ranging in length from 12 nucleotides (corresponding to at least 12 contiguous nucleotides which hybridize under stringent conditions to or are at least 80% identical to a nucleic acid represented by one of SEQ ID Nos. 1-544, preferably SEO ID Nos. 1-168, even more preferably SEO ID Nos. 1-35, or a sequence complementary thereto) up to a maximum length suitable for one or more biological manipulations, including replication and expression, of the nucleic acid molecule. The invention includes but is not limited to (a) nucleic acid having the size of a full gene, and comprising at least one of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto; (b) the nucleic acid of (a) also comprising at least one additional gene, operably linked to permit expression of a fusion protein; (c) an expression vector comprising (a) or (b); (d) a plasmid comprising (a) or (b); and (e) a recombinant viral particle comprising (a) or (b). Construction of (a) can be accomplished as described below in part IV.

The sequence of a nucleic acid of the present invention is not limited and can be any sequence of A, T, G, and/or C (for DNA) and A, U, G, and/or C (for RNA) or modified bases thereof, including inosine and pseudouridine. The choice of sequence will depend on the desired function and can be dictated by coding regions desired, the intron-like regions desired, and the regulatory regions desired.

#### VI. Vectors Carrying Nucleic Acids of the Present Invention

The invention further provides plasmids and vectors, which can be used to express a gene in a host cell. The host cell may be any prokaryotic or eukaryotic cell. Thus, a nucleotide sequence derived from any one of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, encoding all or a selected portion of a protein, can be used to produce a recombinant form of an polypeptide via microbial or eukaryotic cellular processes. Ligating the polynucleotide sequence into a gene construct, such as an expression vector, and transforming or transfecting into hosts, either eukaryotic (yeast, avian, insect or mammalian) or prokaryotic (bacterial cells), are standard procedures well known in the art.

Vectors that allow expression of a nucleic acid in a cell are referred to as expression vectors. Typically, expression vectors contain a nucleic acid operably linked to at least one transcriptional regulatory sequence. Regulatory sequences are art-recognized and are selected to direct expression of the subject nucleic acids. Transcriptional regulatory sequences are described in Goeddel; Gene Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, CA (1990). In one embodiment, the expression vector includes a recombinant gene encoding a peptide having an agonistic activity of a subject polypeptide, or alternatively, encoding a peptide which is an antagonistic form of a subject polypeptide.

The choice of plasmid will depend on the type of cell in which propagation is desired and the purpose of propagation. Certain vectors are useful for amplifying and making large amounts of the desired DNA sequence. Other vectors are suitable for expression in cells in culture. Still other vectors are suitable for transfer and expression in cells in a whole animal or person. The choice of appropriate vector is well within the skill of the art. Many such vectors are available commercially. The nucleic acid or full-length gene is inserted into a vector typically by means of DNA ligase attachment to a cleaved restriction enzyme site in the vector. Alternatively, the desired nucleotide sequence may be inserted by homologous recombination in vivo. Typically this is accomplished by attaching regions of homology to the vector on the flanks of the desired nucleotide sequence. Regions of homology are added by ligation of oligonucleotides, or by polymerase chain reaction using primers comprising both the region of homology and a portion of the desired nucleotide sequence.

Nucleic acids or full-length genes are linked to regulatory sequences as appropriate to obtain the desired expression properties. These may include promoters (attached either at the 5' end of the sense strand or at the 3' end of the antisense strand), enhancers, terminators, operators, repressors, and inducers. The promoters may be regulated or constitutive. In some situations it may be desirable to use conditionally active promoters, such as tissue-specific or developmental stage-specific promoters. These are linked to the desired nucleotide sequence using the techniques described above for linkage to vectors. Any techniques known in the art may be used.

When any of the above host cells, or other appropriate host cells or organisms, are used to replicate and/or express the polynucleotides or nucleic acids of the invention, the resulting replicated nucleic acid, RNA, expressed protein or polypeptide, is within the scope of the invention as a product of the host cell or organism. The product is recovered by any appropriate means known in the art.

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Once the gene corresponding to the nucleic acid is identified, its expression can be regulated in the cell to which the gene is native. For example, an endogenous gene of a cell can be regulated by an exogenous regulatory sequence as disclosed in U.S. Patent No. 5,641,670, "Protein Production and Protein Delivery."

A number of vectors exist for the expression of recombinant proteins in yeast (see, for example, Broach *et al.* (1983) in Experimental Manipulation of Gene Expression, ed. M. Inouye, Academic Press, p. 83, incorporated by reference herein). In addition, drug resistance markers such as ampicillin can be used. In an illustrative embodiment, a polypeptide is produced recombinantly utilizing an expression vector generated by sub-cloning one of the nucleic acids represented in one of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto.

The preferred mammalian expression vectors contain both prokaryotic sequences, to facilitate the propagation of the vector in bacteria, and one or more eukaryotic transcription units that are expressed in eukaryotic cells. The various methods employed in the preparation of plasmids and transformation of host organisms are well known in the art. For other suitable expression systems for both prokaryotic and eukaryotic cells, as well as general recombinant procedures, see Molecular Cloning: A Laboratory Manual, 2<sup>nd</sup> Ed., ed. by Sambrook, Fritsch and Maniatis (Cold Spring Harbor Laboratory Press: 1989) Chapters 16 and 17.

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When it is desirable to express only a portion of a gene, e.g., a truncation mutant, it may be necessary to add a start codon (ATG) to the oligonucleotide fragment containing the desired sequence to be expressed. It is well known in the art that a methionine at the N-terminal position can be enzymatically cleaved by the use of the enzyme methionine aminopeptidase (MAP). MAP has been cloned from E. coli (Ben-Bassat et al. (1987) J. Bacteriol. 169:751-757) and Salmonella typhimurium and its in vitro activity has been demonstrated on recombinant proteins (Miller et al. (1987) PNAS 84:2718-1722). Therefore, removal of an N-terminal methionine, if desired, can be achieved either in vivo by expressing polypeptides in a host which produces MAP (e.g., E. coli or CM89 or S. cerevisiae), or in vitro by use of purified MAP (e.g., procedure of Miller et al., supra).

Moreover, the nucleic acid constructs of the present invention can also be used as part of a gene therapy protocol to deliver nucleic acids such as antisense nucleic acids. Thus, another aspect of the invention features expression vectors for *in vivo* or *in vitro* transfection with an antisense oligonucleotide.

In addition to viral transfer methods, non-viral methods can also be employed to introduce a subject nucleic acid, e.g., a sequence represented by one of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, into the tissue of an animal. Most nonviral methods of gene transfer rely on normal mechanisms used by mammalian cells for the uptake and intracellular transport of macromolecules. In preferred embodiments, non-viral targeting means of the present invention rely on endocytic pathways for the uptake of the subject nucleic acid by the targeted cell. Exemplary targeting means of this type include liposomal derived systems, polylysine conjugates, and artificial viral envelopes.

A nucleic acid of any of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, the corresponding cDNA, or the full-length gene may be used to express the partial or complete gene product. Appropriate nucleic acid constructs are purified using standard recombinant DNA techniques as described in, for example, Sambrook et al., (1989) Molecular Cloning: A Laboratory Manual, 2nd ed. (Cold Spring Harbor Press, Cold Spring Harbor, New York), and under current regulations described in United States Dept. of HHS, National Institute of Health (NIH) Guidelines for Recombinant

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DNA Research. The polypeptides encoded by the nucleic acid may be expressed in any expression system, including, for example, bacterial, yeast, insect, amphibian and mammalian systems. Suitable vectors and host cells are described in U.S. Patent No. 5,654,173.

Bacteria. Expression systems in bacteria include those described in Chang et al., Nature (1978) 275:615, Goeddel et al., Nature (1979) 281:544; Goeddel et al., Nucleic Acids Res. (1980) 8:4057; EP 0 036,776, U.S. Patent No. 4,551,433, DeBoer et al., Proc. Natl. Acad. Sci. (USA) (1983) 80:2125, and Siebenlist et al., Cell (1980) 20:269.

Yeast. Expression systems in yeast include those described in Hinnen et al., 10 Proc. Natl. Acad. Sci. (USA) (1978) 75:1929; Ito et al., J. Bacteriol. (1983) 153:163; Kurtz et al., Mol. Cell. Biol. (1986) 6:142; Kunze et al., J. Basic Microbiol. (1985) 25:141; Gleeson et al., J. Gen. Microbiol. (1986) 132:3459, Roggenkamp et al., Mol. Gen. Genet. (1986) 202:302) Das et al., J. Bacteriol. (1984) 158:1165; De Louvencourt et al., J. Bacteriol. (1983) 154:737, Van den Berg et al., Bio/Technology 15 (1990) 8:135; Kunze et al., J. Basic Microbiol. (1985) 25:141; Cregg et al., Mol. Cell. Biol. (1985) 5:3376, U.S. Patent Nos. 4,837,148 and 4,929,555; Beach and Nurse, Nature (1981) 300:706; Davidow et al., Curr. Genet. (1985) 10:380, Gaillardin et al., Curr. Genet. (1985) 10:49, Ballance et al., Biochem. Biophys. Res. Commun. (1983) 112:284289; Tilburn et al., Gene (1983) 26:205221, Yelton et al., Proc. Natl. Acad. 20 Sci. (USA) (1984) 81:14701474, Kelly and Hynes, EMBO J. (1985) 4:475479; EP 0 244,234, and WO 91/00357.

Insect Cells. Expression of heterologous genes in insects is accomplished as described in U.S. Patent No. 4,745,051, Friesen et al. (1986) "The Regulation of Baculovirus Gene Expression" in: The Molecular Biology Of Baculoviruses (W. Doerfler, ed.), EP 0 127,839, EP 0 155,476, and Vlak et al., J. Gen. Virol. (1988) 69:765776, Miller et al., Ann. Rev. Microbiol. (1988) 42:177, Carbonell et al., Gene (1988) 73:409, Maeda et al., Nature (1985) 315:592594, LebacqVerheyden et al., Mol. Cell. Biol. (1988) 8:3129; Smith et al., Proc. Natl. Acad. Sci. (USA) (1985) 82:8404, Miyajima et al., Gene (1987) 58:273; and Martin et al., DNA (1988) 7:99. Numerous baculoviral strains and variants and corresponding permissive insect host cells from hosts are described in Luckow et al., Bio/Technology (1988) 6:4755, Miller

et al., Generic Engineering (Setlow, J.K. et al. eds.), Vol. 8 (Plenum Publishing, 1986), pp. 277279, and Maeda et al., Nature, (1985) 315:592-594.

Mammalian Cells. Mammalian expression is accomplished as described in Dijkema et al., EMBO J. (1985) 4:761, Gorman et al., Proc. Natl. Acad. Sci. (USA) (1982) 79:6777, Boshart et al., Cell (1985) 41:521 and U.S. Patent No. 4,399,216. Other features of mammalian expression are facilitated as described in Ham and Wallace, Meth. Enz. (1979) 58:44, Barnes and Sato, Anal. Biochem. (1980) 102:255, U.S. Patent Nos. 4,767,704, 4,657,866, 4,927,762, 4,560,655, WO 90/103430, WO 87/00195, and U.S. RE 30,985.

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## VII. Therapeutic Nucleic Acid Constructs

One aspect of the invention relates to the use of the isolated nucleic acid, e.g., SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, in antisense therapy. As used herein, antisense therapy refers to administration or *in situ* generation of oligonucleotide molecules or their derivatives which specifically hybridize (e.g., bind) under cellular conditions with the cellular mRNA and/or genomic DNA, thereby inhibiting transcription and/or translation of that gene. The binding may be by conventional base pair complementarity, or, for example, in the case of binding to DNA duplexes, through specific interactions in the major groove of the double helix. In general, antisense therapy refers to the range of techniques generally employed in the art, and includes any therapy which relies on specific binding to oligonucleotide sequences.

An antisense construct of the present invention can be delivered, for example, as an expression plasmid which, when transcribed in the cell, produces RNA which is complementary to at least a unique portion of the cellular mRNA. Alternatively, the antisense construct is an oligonucleotide probe which is generated *ex vivo* and which, when introduced into the cell, causes inhibition of expression by hybridizing with the mRNA and/or genomic sequences of a subject nucleic acid. Such oligonucleotide probes are preferably modified oligonucleotides which are resistant to endogenous nucleases, e.g., exonucleases and/or endonucleases, and are therefore stable *in vivo*. Exemplary nucleic acid molecules for use as antisense oligonucleotides are phosphoramidate, phosphorothioate and methylphosphonate analogs of DNA (see also

U.S. Patents 5,176,996; 5,264,564; and 5,256,775). Additionally, general approaches to constructing oligomers useful in antisense therapy have been reviewed, for example, by Van der Krol et al. (1988) BioTechniques 6:958-976; and Stein et al. (1988) Cancer Res 48:2659-2668. With respect to antisense DNA, oligodeoxyribonucleotides derived from the translation initiation site, e.g., between the -10 and +10 regions of the nucleotide sequence of interest, are preferred.

Antisense approaches involve the design of oligonucleotides (either DNA or RNA) that are complementary to mRNA. The antisense oligonucleotides will bind to the mRNA transcripts and prevent translation. Absolute complementarity, although preferred, is not required. In the case of double-stranded antisense nucleic acids, a single strand of the duplex DNA may thus be tested, or triplex formation may be assayed. The ability to hybridize will depend on both the degree of complementarity and the length of the antisense nucleic acid. Generally, the longer the hybridizing nucleic acid, the more base mismatches with an RNA it may contain and still form a stable duplex (or triplex, as the case may be). One skilled in the art can ascertain a tolerable degree of mismatch by use of standard procedures to determine the melting point of the hybridized complex.

Oligonucleotides that are complementary to the 5' end of the mRNA, e.g., the 5' untranslated sequence up to and including the AUG initiation codon, should work most efficiently at inhibiting translation. However, sequences complementary to the 3' untranslated sequences of mRNAs have recently been shown to be effective at inhibiting translation of mRNAs as well. (Wagner, R. 1994. Nature 372:333). Therefore, oligonucleotides complementary to either the 5' or 3' untranslated, noncoding regions of a gene could be used in an antisense approach to inhibit translation of endogenous mRNA. Oligonucleotides complementary to the 5' untranslated region of the mRNA should include the complement of the AUG start codon. Antisense oligonucleotides complementary to mRNA coding regions are typically less efficient inhibitors of translation but could also be used in accordance with the invention. Whether designed to hybridize to the 5', 3', or coding region of subject mRNA, antisense nucleic acids should be at least six nucleotides in length, and are preferably less that about 100 and more preferably less than about 50, 25, 17 or 10 nucleotides in length.

Regardless of the choice of target sequence, it is preferred that *in vitro* studies are first performed to quantitate the ability of the antisense oligonucleotide to quantitate the ability of the antisense oligonucleotide to inhibit gene expression. It is preferred that these studies utilize controls that distinguish between antisense gene inhibition and nonspecific biological effects of oligonucleotides. It is also preferred that these studies compare levels of the target RNA or protein with that of an internal control RNA or protein. Additionally, it is envisioned that results obtained using the antisense oligonucleotide are compared with those obtained using a control oligonucleotide. It is preferred that the control oligonucleotide is of approximately the same length as the test oligonucleotide and that the nucleotide sequence of the oligonucleotide differs from the antisense sequence no more than is necessary to prevent specific hybridization to the target sequence.

The oligonucleotides can be DNA or RNA or chimeric mixtures or derivatives or modified versions thereof, single-stranded or double-stranded. The oligonucleotide can be modified at the base moiety, sugar moiety, or phosphate backbone, for example, to improve stability of the molecule, hybridization, etc. The oligonucleotide may include other appended groups such as peptides (e.g., for targeting host cell receptors), or agents facilitating transport across the cell membrane (see, e.g., Letsinger et al., 1989, Proc. Natl. Acad. Sci. U.S.A. 86:6553-6556; Lemaitre et al., 1987, Proc. Natl. Acad. Sci. 84:648-652; PCT Publication No. WO 88/09810, published December 15, 1988) or the blood-brain barrier (see, e.g., PCT Publication No. WO 89/10134, published April 25, 1988), hybridization-triggered cleavage agents (See, e.g., Krol et al., 1988, BioTechniques 6:958-976), or intercalating agents (See, e.g., Zon, 1988, Pharm. Res. 5:539-549). To this end, the oligonucleotide may be conjugated to another molecule, e.g., a peptide, hybridization triggered cross-linking agent, transport agent, hybridization-triggered cleavage agent, etc.

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The antisense oligonucleotide may comprise at least one modified base moiety which is selected from the group including but not limited to 5-fluorouracil, 5-bromouracil, 5-chlorouracil, 5-iodouracil, hypoxanthine, xantine, 4-acetylcytosine, 5-(carboxyhydroxytriethyl) uracil, 5-carboxymethylaminomethyl-2-thiouridine, 5-carboxymethylaminomethyluracil, dihydrouracil, beta-D-galactosylqueosine, inosine, N6-isopentenyladenine, 1-methylguanine, 1-methylinosine, 2,2-dimethylguanine, 2-methylguanine, 3-methylcytosine, 5-methylcytosine, N6-adenine,

7-methylguanine, 5-methylaminomethyluracil, 5-methoxyaminomethyl-2-thiouracil, beta-D-mannosylqueosine, 5-methoxycarboxymethyluracil, 5-methoxyuracil, 2-methylthio-N6-isopentenyladenine, uracil-5-oxyacetic acid (v), wybutoxosine, pseudouracil, queosine, 2-thiocytosine, 5-methyl-2-thiouracil, 2-thiouracil, 4-thiouracil, 5-methyluracil, uracil-5-oxyacetic acid methylester, uracil-5-oxyacetic acid (v), 5-methyl-2-thiouracil, 3-(3-amino-3-N-2-carboxypropyl) uracil, (acp3)w, and 2.6-diaminopurine.

The antisense oligonucleotide may also comprise at least one modified sugar moiety selected from the group including but not limited to arabinose, 2-fluoroarabinose, xylulose, and hexose.

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The antisense oligonucleotide can also contain a neutral peptide-like backbone. Such molecules are termed peptide nucleic acid (PNA)-oligomers and are described, e.g., in Perry- O'Keefe et al. (1996) Proc. Natl. Acad. Sci. U.S.A. 93:14670 and in Eglom et al. (1993) Nature 365:566. One advantage of PNA oligomers is their capability to bind to complementary DNA essentially independently from the ionic strength of the medium due to the neutral backbone of the DNA. In yet another embodiment, the antisense oligonucleotide comprises at least one modified phosphate backbone selected from the group consisting of a phosphorothioate, a phosphorodithioate, a phosphoramidothioate, a phosphoramidate, a methylphosphonate, an alkyl phosphotriester, and a formacetal or analog thereof.

In yet a further embodiment, the antisense oligonucleotide is an α-anomeric oligonucleotide. An α-anomeric oligonucleotide forms specific double-stranded hybrids with complementary RNA in which, contrary to the usual β-units, the strands run parallel to each other (Gautier et al., 1987, Nucl. Acids Res. 15:6625-6641). The oligonucleotide is a 2'-O-methylribonucleotide (Inoue et al., 1987, Nucl. Acids Res. 15:6131-12148), or a chimeric RNA-DNA analogue (Inoue et al., 1987, FEBS Lett. 215:327-330).

Oligonucleotides of the invention may be synthesized by standard methods known in the art, e.g., by use of an automated DNA synthesizer (such as are commercially available from Biosearch, Applied Biosystems, etc.). As examples, phosphorothicate oligonucleotides may be synthesized by the method of Stein et al. (1988, Nucl. Acids Res. 16:3209), methylphosphonate olgonucleotides can be

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prepared by use of controlled pore glass polymer supports (Sarin et al., 1988, Proc. Natl. Acad. Sci. U.S.A. 85:7448-7451), etc.

While antisense nucleotides complementary to a coding region sequence can be used, those complementary to the transcribed untranslated region and to the region comprising the initiating methionine are most preferred.

The antisense molecules can be delivered to cells which express the target nucleic acid *in vivo*. A number of methods have been developed for delivering antisense DNA or RNA to cells; e.g., antisense molecules can be injected directly into the tissue site, or modified antisense molecules, designed to target the desired cells (e.g., antisense linked to peptides or antibodies that specifically bind receptors or antigens expressed on the target cell surface) can be administered systemically.

However, it is often difficult to achieve intracellular concentrations of the antisense sufficient to suppress translation on endogenous mRNAs. Therefore, a preferred approach utilizes a recombinant DNA construct in which the antisense oligonucleotide is placed under the control of a strong pol III or pol II promoter. The use of such a construct to transfect target cells in the patient will result in the transcription of sufficient amounts of single stranded RNAs that will form complementary base pairs with the endogenous transcripts and thereby prevent translation of the target mRNA. For example, a vector can be introduced in vivo such that it is taken up by a cell and directs the transcription of an antisense RNA. Such a vector can remain episomal or become chromosomally integrated, as long as it can be transcribed to produce the desired antisense RNA. Such vectors can be constructed by recombinant DNA technology methods standard in the art. Vectors can be plasmid, viral, or others known in the art for replication and expression in mammalian cells. Expression of the sequence encoding the antisense RNA can be by any promoter known in the art to act in mammalian, preferably human cells. Such promoters can be inducible or constitutive. Such promoters include but are not limited to: the SV40 early promoter region (Bernoist and Chambon, 1981, Nature 290:304-310), the promoter contained in the 3' long terminal repeat of Rous sarcoma virus (Yamamoto et al., 1980, Cell 22:787-797), the herpes thymidine kinase promoter (Wagner et al., 1981, Proc. Natl. Acad. Sci. U.S.A. 78:1441-1445), the regulatory sequences of the metallothionein gene (Brinster et al, 1982, Nature 296:39-42), etc. Any type of plasmid, cosmid, YAC or viral vector can be used to prepare the recombinant DNA

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construct which can be introduced directly into the tissue site; e.g., the choroid plexus or hypothalamus. Alternatively, viral vectors can be used which selectively infect the desired tissue (e.g., for brain, herpesvirus vectors may be used), in which case administration may be accomplished by another route (e.g., systemically).

In another aspect of the invention, ribozyme molecules designed to catalytically cleave target mRNA transcripts can be used to prevent translation of target mRNA and expression of a target protein (See, e.g., PCT International Publication WO90/11364, published October 4, 1990; Sarver et al., 1990, Science 247:1222-1225 and U.S. Patent No. 5,093,246). While ribozymes that cleave mRNA at site specific recognition sequences can be used to destroy target mRNAs, the use of hammerhead ribozymes is preferred. Hammerhead ribozymes cleave mRNAs at locations dictated by flanking regions that form complementary base pairs with the target mRNA. The sole requirement is that the target mRNA have the following sequence of two bases: 5'-UG-3'. The construction and production of hammerhead ribozymes is well known in the art and is described more fully in Haseloff and Gerlach, 1988, Nature, 334:585-591. Preferably the ribozyme is engineered so that the cleavage recognition site is located near the 5' end of the target mRNA; i.e., to increase efficiency and minimize the intracellular accumulation of non-functional mRNA transcripts.

The ribozymes of the present invention also include RNA endoribonucleases (hereinafter "Cech-type ribozymes") such as the one which occurs naturally in *Tetrahymena thermophila* (known as the IVS, or L-19 IVS RNA) and which has been extensively described by Thomas Cech and collaborators (Zaug, et al., 1984, Science, 224:574-578; Zaug and Cech, 1986, Science, 231:470-475; Zaug, et al., 1986, Nature, 324:429-433; published International patent application No. WO88/04300 by University Patents Inc.; Been and Cech, 1986, Cell, 47:207-216). The Cech-type ribozymes have an eight base pair active site which hybridizes to a target RNA sequence whereafter cleavage of the target RNA takes place. The invention encompasses those Cech-type ribozymes which target eight base-pair active site sequences that are present in a target gene.

As in the antisense approach, the ribozymes can be composed of modified oligonucleotides (e.g., for improved stability, targeting, etc.) and should be delivered to cells which express the target gene *in vivo*. A preferred method of delivery

involves using a DNA construct "encoding" the ribozyme under the control of a strong constitutive pol III or pol II promoter, so that transfected cells will produce sufficient quantities of the ribozyme to destroy endogenous messages and inhibit translation. Because ribozymes, unlike antisense molecules, are catalytic, a lower intracellular concentration is required for efficiency.

Antisense RNA, DNA, and ribozyme molecules of the invention may be prepared by any method known in the art for the synthesis of DNA and RNA molecules. These include techniques for chemically synthesizing oligodeoxyribonucleotides and oligoribonucleotides well known in the art such as for example solid phase phosphoramidite chemical synthesis. Alternatively, RNA molecules may be generated by *in vitro* and *in vivo* transcription of DNA sequences encoding the antisense RNA molecule. Such DNA sequences may be incorporated into a wide variety of vectors which incorporate suitable RNA polymerase promoters such as the T7 or SP6 polymerase promoters. Alternatively, antisense cDNA constructs that synthesize antisense RNA constitutively or inducibly, depending on the promoter used, can be introduced stably into cell lines.

Moreover, various well-known modifications to nucleic acid molecules may be introduced as a means of increasing intracellular stability and half-life. Possible modifications include but are not limited to the addition of flanking sequences of ribonucleotides or deoxyribonucleotides to the 5' and/or 3' ends of the molecule or the use of phosphorothioate or 2' O-methyl rather than phosphodiesterase linkages within the oligodeoxyribonucleotide backbone.

## VIII. Polypeptides of the Present Invention

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The present invention makes available isolated polypeptides which are isolated from, or otherwise substantially free of other cellular proteins, especially other signal transduction factors and/or transcription factors which may normally be associated with the polypeptide. Subject polypeptides of the present invention include polypeptides encoded by the nucleic acids of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, or polypeptides encoded by genes of which a sequence in SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, is a fragment. Polypeptides of the present invention

include those proteins which are differentially regulated in tumor cells, especially colon cancer-derived cell lines (relative to normal cells, e.g., normal colon tissue and non-colon tissue). In preferred embodiments, the polypeptides are upregulated in tumor cells, especially colon cancer cancer-derived cell lines. In other embodiments, the polypeptides are downregulated in tumor cells, especially colon cancer-derived cell lines. Proteins which are upregulated, such as oncogenes, or downregulated, such as tumor suppressors, in aberrantly proliferating cells may be targets for diagnostic or therapeutic techniques. For example, upregulation of the *cdc2* gene induces mitosis. Overexpression of the *myt1* gene, a mitotic deactivator, negatively regulates the activity of *cdc2*. Aberrant proliferation may thus be induced either by upregulating *cdc2* or by downregulating *myt1*.

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The term "substantially free of other cellular proteins" (also referred to herein as "contaminating proteins") or "substantially pure or purified preparations" are defined as encompassing preparations of polypeptides having less than about 20% (by dry weight) contaminating protein, and preferably having less than about 5% contaminating protein. Functional forms of the subject polypeptides can be prepared, for the first time, as purified preparations by using a cloned nucleic acid as described herein. Full length proteins or fragments corresponding to one or more particular motifs and/or domains or to arbitrary sizes, for example, at least about 5, 10, 25, 50, 75, or 100 amino acids in length are within the scope of the present invention.

For example, isolated polypeptides can be encoded by all or a portion of a nucleic acid sequence shown in any of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto. Isolated peptidyl portions of proteins can be obtained by screening peptides recombinantly produced from the corresponding fragment of the nucleic acid encoding such peptides. In addition, fragments can be chemically synthesized using techniques known in the art such as conventional Merrifield solid phase f-Moc or t-Boc chemistry. For example, a polypeptide of the present invention may be arbitrarily divided into fragments of desired length with no overlap of the fragments, or preferably divided into overlapping fragments of a desired length. The fragments can be produced (recombinantly or by chemical synthesis) and tested to identify those peptidyl fragments which can function as either agonists or antagonists of a wild-type (e.g., "authentic") protein.

Another aspect of the present invention concerns recombinant forms of the subject proteins. Recombinant polypeptides preferred by the present invention, in addition to native proteins, as described above are encoded by a nucleic acid, which is at least 60%, more preferably at least 80%, and more preferably 85%, and more preferably 90%, and more preferably 95% identical to an amino acid sequence encoded by SEQ ID Nos. 1-544. Polypeptides which are encoded by a nucleic acid that is at least about 98-99% identical with the sequence of SEQ ID Nos. 1-544 are also within the scope of the invention. Also included in the present invention are peptide fragments comprising at least a portion of such a protein.

In a preferred embodiment, a polypeptide of the present invention is a mammalian polypeptide and even more preferably a human polypeptide. In particularly preferred embodiment, the polypeptide retains wild-type bioactivity. It will be understood that certain post-translational modifications, e.g., phosphorylation and the like, can increase the apparent molecular weight of the polypeptide relative to the unmodified polypeptide chain.

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The present invention further pertains to recombinant forms of one of the subject polypeptides. Such recombinant polypeptides preferably are capable of functioning in one of either role of an agonist or antagonist of at least one biological activity of a wild-type ("authentic") polypeptide of the appended sequence listing. The term "evolutionarily related to", with respect to amino acid sequences of proteins, refers to both polypeptides having amino acid sequences which have arisen naturally, and also to mutational variants of human polypeptides which are derived, for example, by combinatorial mutagenesis.

In general, polypeptides referred to herein as having an activity (e.g., are "bioactive") of a protein are defined as polypeptides which include an amino acid sequence encoded by all or a portion of the nucleic acid sequences shown in one of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, and which mimic or antagonize all or a portion of the biological/biochemical activities of a naturally occurring protein. According to the present invention, a polypeptide has biological activity if it is a specific agonist or antagonist of a naturally occurring form of a protein.

Assays for determining whether a compound, e.g, a protein or variant thereof, has one or more of the above biological activities are well known in the art. In certain

embodiments, the polypeptides of the present invention have activities such as those outlined above.

In another embodiment, the coding sequences for the polypeptide can be incorporated as a part of a fusion gene including a nucleotide sequence encoding a different polypeptide. This type of expression system can be useful under conditions where it is desirable to produce an immunogenic fragment of a polypeptide (see, for example, EP Publication No: 0259149; and Evans et al. (1989) Nature 339:385; Huang et al. (1988) J. Virol. 62:3855; and Schlienger et al. (1992) J. Virol. 66:2). In addition to utilizing fusion proteins to enhance immunogenicity, it is widely appreciated that fusion proteins can also facilitate the expression of proteins, and, accordingly, can be used in the expression of the polypeptides of the present invention (see, for example, Current Protocols in Molecular Biology, eds. Ausubel et al. (N.Y.: John Wiley & Sons, 1991)). In another embodiment, a fusion gene coding for a purification leader sequence, such as a poly-(His)/enterokinase cleavage site sequence at the N-terminus of the desired portion of the recombinant protein, can allow purification of the expressed fusion protein by affinity chromatography using a Ni<sup>2+</sup> metal resin. The purification leader sequence can then be subsequently removed by treatment with enterokinase to provide the purified protein (e.g., see Hochuli et al. (1987) J. Chromatography 411:177; and Janknecht et al. PNAS 88:8972).

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Techniques for making fusion genes are known to those skilled in the art. Essentially, the joining of various DNA fragments coding for different polypeptide sequences is performed in accordance with conventional techniques, employing blunt-ended or stagger-ended termini for ligation, restriction enzyme digestion to provide for appropriate termini, filling-in of cohesive ends as appropriate, alkaline phosphatase treatment to avoid undesirable joining, and enzymatic ligation. In another embodiment, the fusion gene can be synthesized by conventional techniques including automated DNA synthesizers. Alternatively, PCR amplification of nucleic acid fragments can be carried out using anchor primers which give rise to complementary overhangs between two consecutive nucleic acid fragments which can subsequently be annealed to generate a chimeric nucleic acid sequence (see, for example, Current Protocols in Molecular Biology, eds. Ausubel et al. John Wiley & Sons: 1992).

The present invention further pertains to methods of producing the subject polypeptides. For example, a host cell transfected with a nucleic acid vector directing

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expression of a nucleotide sequence encoding the subject polypeptides can be cultured under appropriate conditions to allow expression of the peptide to occur. Suitable media for cell culture are well known in the art. The recombinant polypeptide can be isolated from cell culture medium, host cells, or both using techniques known in the art for purifying proteins including ion-exchange chromatography, gel filtration chromatography, ultrafiltration, electrophoresis, and immunoaffinity purification with antibodies specific for such peptide. In a preferred embodiment, the recombinant polypeptide is a fusion protein containing a domain which facilitates its purification, such as GST fusion protein.

Moreover, it will be generally appreciated that, under certain circumstances, it may be advantageous to provide homologs of one of the subject polypeptides which function in a limited capacity as one of either an agonist (mimetic) or an antagonist, in order to promote or inhibit only a subset of the biological activities of the naturally occurring form of the protein. Thus, specific biological effects can be elicited by treatment with a homolog of limited function, and with fewer side effects relative to treatment with agonists or antagonists which are directed to all of the biological activities of naturally occurring forms of subject proteins.

Homologs of each of the subject polypeptide can be generated by mutagenesis, such as by discrete point mutation(s), or by truncation. For instance, mutation can give rise to homologs which retain substantially the same, or merely a subset, of the biological activity of the polypeptide from which it was derived. Alternatively, antagonistic forms of the polypeptide can be generated which are able to inhibit the function of the naturally occurring form of the protein, such as by competitively binding to a receptor.

The recombinant polypeptides of the present invention also include homologs of the wild-type proteins, such as versions of those proteins which are resistant to proteolytic cleavage, for example, due to mutations which alter ubiquitination or other enzymatic targeting associated with the protein.

Polypeptides may also be chemically modified to create derivatives by forming covalent or aggregate conjugates with other chemical moieties, such as glycosyl groups, lipids, phosphate, acetyl groups and the like. Covalent derivatives of proteins can be prepared by linking the chemical moieties to functional groups on

amino acid sidechains of the protein or at the N-terminus or at the C-terminus of the polypeptide.

Modification of the structure of the subject polypeptides can be for such purposes as enhancing therapeutic or prophylactic efficacy, stability (e.g., ex vivo shelf life and resistance to proteolytic degradation), or post-translational modifications (e.g., to alter phosphorylation pattern of protein). Such modified peptides, when designed to retain at least one activity of the naturally occurring form of the protein, or to produce specific antagonists thereof, are considered functional equivalents of the polypeptides described in more detail herein. Such modified peptides can be produced, for instance, by amino acid substitution, deletion, or addition. The substitutional variant may be a substituted conserved amino acid or a substituted nonconserved amino acid.

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For example, it is reasonable to expect that an isolated replacement of a leucine with an isoleucine or valine, an aspartate with a glutamate, a threonine with a serine, or a similar replacement of an amino acid with a structurally related amino acid (i.e., isosteric and/or isoelectric mutations) will not have a major effect on the biological activity of the resulting molecule. Conservative replacements are those that take place within a family of amino acids that are related in their side chains. Genetically encoded amino acids can be divided into four families: (1) acidic = aspartate, glutamate; (2) basic = lysine, arginine, histidine; (3) nonpolar = alanine, valine, leucine, isoleucine, proline, phenylalanine, methionine, tryptophan; and (4) uncharged polar = glycine, asparagine, glutamine, cysteine, serine, threonine, tyrosine. In similar fashion, the amino acid repertoire can be grouped as (1) acidic = aspartate, glutamate; (2) basic = lysine, arginine histidine, (3) aliphatic = glycine, alanine, valine, leucine, isoleucine, serine, threonine, with serine and threonine optionally be grouped separately as aliphatic-hydroxyl; (4) aromatic = phenylalanine, tyrosine, tryptophan; (5) amide = asparagine, glutamine; and (6) sulfur -containing = cysteine and methionine. (see, for example, Biochemistry, 2<sup>nd</sup> ed., Ed. by L. Stryer, WH Freeman and Co.: 1981). Whether a change in the amino acid sequence of a peptide results in a functional homolog (e.g., functional in the sense that the resulting polypeptide mimics or antagonizes the wild-type form) can be readily determined by assessing the ability of the variant peptide to produce a response in cells in a fashion similar to the wild-type protein, or competitively inhibit such a response.

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Polypeptides in which more than one replacement has taken place can readily be tested in the same manner. The variant may be designed so as to retain biological activity of a particular region of the protein. In a non-limiting example, Osawa et al., 1994, Biochemistry and Molecular International 34:1003-1009, discusses the actin binding region of a protein from several different species. The actin binding regions of the these species are considered homologous based on the fact that they have amino acids that fall within "homologous residue groups." Homologous residues are judged according to the following groups (using single letter amino acid designations): STAG; ILVMF; HRK; DEQN; and FYW. For example, an S, a T, an A or a G can be in a position and the function (in this case actin binding) is retained.

Additional guidance on amino acid substitution is available from studies of protein evolution. Go et al., 1980, Int. J. Peptide Protein Res. 15:211-224, classified amino acid residue sites as interior or exterior depending on their accessibility. More frequent substitution on exterior sites was confirmed to be general in eight sets of homologous protein families regardless of their biological functions and the presence or absence of a prosthetic group. Virtually all types of amino acid residues had higher mutabilities on the exterior than in the interior. No correlation between mutability and polarity was observed of amino acid residues in the interior and exterior, respectively. Amino acid residues were classified into one of three groups depending on their polarity: polar (Arg, Lys, His, Gln, Asn, Asp, and Glu); weak polar (Ala, Pro, Gly, Thr, and Ser), and nonpolar (Cys, Val, Met, Ile, Leu, Phe, Tyr, and Trp). Amino acid replacements during protein evolution were very conservative: 88% and 76% of them in the interior or exterior, respectively, were within the same group of the three. Intergroup replacements are such that weak polar residues are replaced more often by nonpolar residues in the interior and more often by polar residues on the exterior.

Querol et al., 1996, Prot. Eng. 9:265-271, provides general rules for amino acid substitutions to enhance protein thermostability. New glycosylation sites can be introduced as discussed in Olsen and Thomsen, 1991, J. Gen. Microbiol. 137:579-585. An additional disulfide bridge can be introduced, as discussed by Perry and Wetzel, 1984, Science 226:555-557; Pantoliano et al., 1987, Biochemistry 26:2077-2082; Matsumura et al., 1989, Nature 342:291-293; Nishikawa et al., 1990, Protein Eng. 3:443-448; Takagi et al., 1990, J. Biol. Chem. 265:6874-6878; Clarke et al., 1993, Biochemistry 32:4322-4329; and Wakarchuk et al., 1994, Protein Eng. 7:1379-1386.

An additional metal binding site can be introduced, according to Toma et al., 1991, Biochemistry 30:97-106, and Haezerbrouck et al., 1993, Protein Eng. 6:643-649. Substitutions with prolines in loops can be made according to Masul et al., 1994, Appl. Env. Microbiol. 60:3579-3584; and Hardy et al., FEBS Lett. 317:89-92.

Cysteine-depleted muteins are considered variants within the scope of the invention. These variants can be constructed according to methods disclosed in U.S. Patent No. 4,959,314, which discloses how to substitute other amino acids for cysteines, and how to determine biological activity and effect of the substitution. Such methods are suitable for proteins according to this invention that have cysteine residues suitable for such substitutions, for example to eliminate disulfide bond formation.

To learn the identity and function of the gene that correlates with an nucleic acid, the nucleic acids or corresponding amino acid sequences can be screened against profiles of protein families. Such profiles focus on common structural motifs among proteins of each family. Publicly available profiles are described above. Additional or alternative profiles are described below.

In comparing a new nucleic acid with known sequences, several alignment tools are available. Examples include PileUp, which creates a multiple sequence alignment, and is described in Feng et al., J. Mol. Evol. (1987) 25:351-360. Another method, GAP, uses the alignment method of Needleman et al., J. Mol. Biol. (1970) 48:443-453. GAP is best suited for global alignment of sequences. A third method, BestFit, functions by inserting gaps to maximize the number of matches using the local homology algorithm of Smith and Waterman, Adv. Appl. Math. (1981) 2:482-489.

Examples of such profiles are described below.

#### **Chemokines**

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Chemokines are a family of proteins that have been implicated in lymphocyte trafficking, inflammatory diseases, angiogenesis, hematopoiesis, and viral infection. See, for example, Rollins, *Blood* (1997) 90(3):909-928, and Wells et al., J. Leuk. Biol. (1997) 61:545-550. U.S. Patent No. 5,605,817 discloses DNA encoding a chemokine expressed in fetal spleen. U.S. Patent No. 5,656,724 discloses chemokine-like

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proteins and methods of use. U.S. Patent No. 5,602,008 discloses DNA encoding a chemokine expressed by liver.

Mutants of the encoded chemokines are polypeptides having an amino acid sequence that possesses at least one amino acid substitution, addition, or deletion as compared to native chemokines. Fragments possess the same amino acid sequence of the native chemokines; mutants may lack the amino and/or carboxyl terminal sequences. Fusions are mutants, fragments, or the native chemokines that also include amino and/or carboxyl terminal amino acid extensions.

The number or type of the amino acid changes is not critical, nor is the length or number of the amino acid deletions, or amino acid extensions that are incorporated in the chemokines as compared to the native chemokine amino acid sequences. A polynucleotide encoding one of these variant polypeptides will retain at least about 80% amino acid identity with at least one known chemokine. Preferably, these polypeptides will retain at least about 85% amino acid sequence identity, more preferably, at least about 90%; even more preferably, at least about 95%. In addition, the variants will exhibit at least 80%; preferably about 90%; more preferably about 95% of at least one activity exhibited by a native chemokine. Chemokine activity includes immunological, biological, receptor binding, and signal transduction functions of the native chemokine.

Chemotaxis. Assays for chemotaxis relating to neutrophils are described in Walz et al., Biochem. Biophys. Res. Commun. (1987) 149:755, Yoshimura et al., Proc. Natl. Acad. Sci. (USA) (1987) 84:9233, and Schroder et al., J. Immunol. (1987) 139:3474; to lymphocytes, Larsen et al., Science (1989) 243:1464, Carr et al., Proc. Natl. Acad. Sci. (USA) (1994) 91:3652; to tumor-infiltrating lymphocytes, Liao et al., J. Exp. Med (1995). 182:1301; to hemopoietic progenitors, Aiuti et al., J. Exp. Med. (1997) 185:111; to monocytes, Valente et al., Biochem. (1988) 27:4162; and to natural killer cells, Loetscher et al., J. Immunol. (1996) 156:322, and Allavena et al., Eur. J. Immunol. (1994) 24:3233.

Assays for determining the biological activity of attracting eosinophils are described in Dahinden et al., J. Exp. Med. (1994) 179:751, Weber et al., J. Immunol. (1995) 154:4166, and Noso et al., Biochem. Biophys. Res. Commun. (1994) 200:1470; for attracting dendritic cells, Sozzani et al., J. Immunol. (1995) 155:3292; for attracting basophils, in Dahinden et al., J. Exp. Med. (1994) 179:751, Alam et al., J.

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Immunol. (1994) 152:1298, Alam et al., J. Exp. Med. (1992) 176:781; and for activating neutrophils, Maghazaci et al., Eur. J. Immunol. (1996) 26:315, and Taub et al., J. Immunol. (1995) 155:3877. Native chemokines can act as mitogens for fibroblasts, assayed as described in Mullenbach et al., J. Biol. Chem. (1986) 261:719.

Receptor Binding. Native chemokines exhibit binding activity with a number of receptors. Description of such receptors and assays to detect binding are described in, for example, Murphy et al., Science (1991) 253:1280; Combadiere et al., J. Biol. Chem. (1995) 270:29671; Daugherty et al., J. Exp. Med. (1996) 183:2349; Samson et al., Biochem. (1996) 35:3362; Raport et al., J. Biol. Chem. (1996) 271:17161; Combadiere et al., J. Leukoc. Biol. (1996) 60:147; Baba et al., J. Biol. Chem. (1997) 23:14893; Yosida et al., J. Biol. Chem. (1997) 272:13803; Arvannitakis et al., Nature (1997) 385:347, and many other assays are known in the art.

Kinase Activiation. Assays for kinase activation are described by Yen et al., J. Leukoc. Biol. (1997) 61:529; Dubois et al., J. Immunol. (1996) 156:1356; Turner et al., J. Immunol. (1995) 155:2437. Assays for inhibition of angiogenesis or cell proliferation are described in Maione et al., Science (1990) 247:77.

Glycosaminoglycan production can be induced by native chemokines, assayed as described in Castor et al., Proc. Natl. Acad. Sci. (USA) (1983) 80:765. Chemokinemediated histamine release from basophils is assayed as described in Dahinden et al., J. Exp. Med. (1989) 170:1787; and White et al., Immunol. Lett. (1989) 22:151.

Heparin binding is described in Luster et al., J. Exp. Med. (1995) 182:219.

Dimerization Activity. Chemokines can possess dimerization activity, which can be assayed according to Burrows et al., Biochem. (1994) 33:12741; and Zhang et al., Mol. Cell. Biol. (1995) 15:4851. Native chemokines can play a role in the inflammatory response of viruses. This activity can be assayed as described in Bleul et al., Nature (1996) 382:829; and Oberlin et al., Nature (1996) 382:833. Exocytosis of monocytes can be promoted by native chemokines. The assay for such activity is described in Uguccioni et al., Eur. J. Immunol. (1995) 25:64. Native chemokines also can inhibit hemapoietic stem cell proliferation. The method for testing for such activity is reported in Graham et al., Nature (1990) 344:442.

<u>Death Domain Proteins</u> Several protein families contain death domain motifs (Feinstein and Kimchi, *TIBS Letters* (1995) 20:242-244). Some death domain-containing proteins are implicated in cytotoxic intracellular signaling (Cleveland and

Ihle, Cell (1995) 81:479-482, Pan et al, Science (1997) 276:111-113, Duan and Dixit, Nature (1997) 385:86-89, and Chinnaiyan et al, Science (1996) 274:990-992). U.S. Patent No. 5,563,039 describes a protein homologous to TRADD (Tumor Necrosis Factor Receptor-1 Associated Death Domain containing protein), and modifications of the active domain of TRADD that retain the functional characteristics of the protein, as well as apoptosis assays for testing the function of such death domain containing proteins. U.S. Patent No. 5,658,883 discloses biologically active TGF-B1 peptides. U.S. Patent No. 5,674,734 discloses protein RIP which contains a C-terminal death domain and an N-terminal kinase domain.

Leukemia Inhibitory Factor (LIF) An LIF profile is constructed from sequences of leukemia inhibitor factor, CT-1 (cardiotrophin-1), CNTF (ciliary neurotrophic factor), OSM (oncostatin M), and IL-6 (interleukin-6). This profile encompasses a family of secreted cytokines that have pleiotropic effects on many cell types including hepatocytes, osteoclasts, neuronal cells and cardiac myocytes, and can be used to detect additional genes encoding such proteins. These molecules are all structurally related and share a common co-receptor gp130 which mediates intracellular signal transduction by cytoplasmic tyrosine kinases such as src.

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Novel proteins related to this family are also likely to be secreted, to activate gp130 and to function in the development of a variety of cell types. Thus new members of this family would be candidates to be developed as growth or survival factors for the cell types that they stimulate. For more details on this family of cytokines, see Pennica et al, Cytokine and Growth Factor Reviews (1996) 7:81-91. U.S. Patent No. 5,420,247 discloses LIF receptor and fusion proteins. U.S. Patent No. 5,443,825 discloses human LIF.

Angiopoietin Angiopoietin-1 is a secreted ligand of the TIE-2 tyrosine kinase; it functions as an angiogenic factor critical for normal vascular development. Angiopoietin-2 is a natural antagonist of angiopoietin-1 and thus functions as an anti-angiogenic factor. These two proteins are structurally similar and activate the same receptor. (Folkman and D'Amore, Cell (1996) 87:1153-1155, and Davis et al., Cell (1996) 87:1161-1169.)

The angiopoietin molecules are composed of two domains, a coiled-coil region and a region related to fibrinogen. The fibrinogen domain is found in many molecules including ficolin and tesascin, and is well defined structurally with many members.

Receptor Protein-Tyrosine Kinases Receptor Protein-Tyrosine Kinases or RPTKs are described in Lindberg, Annu. Rev. Cell Biol. (1994) 10:251-337.

Growth Factors: Epidermal Growth Factor (EGF) and Fibroblast Growth Factor (FGF)

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For a discussion of growth factor superfamilies, see <u>Growth Factors</u>: <u>A Practical Approach</u>, Appendix A1 (Ed. McKay and Leigh, Oxford University Press, NY, 1993) pp. 237-243.

The alignments (pretty box) for EGF and FGF are shown in Figures 1 and 2, respectively. U.S. Patent No. 4,444,760 discloses acidic brain fibroblast growth factor, which is active in the promotion of cell division and wound healing. U.S. Patent No. 5,439,818 discloses DNA encoding human recombinant basic fibroblast growth factor, which is active in wound healing. U.S. Patent No. 5,604,293 discloses recombinant human basic fibroblast growth factor, which is useful for wound healing. U.S. Patent No. 5,410,832 discloses brain-derived and recombinant acidic fibroblast growth factor, which act as mitogens for mesoderm and neuroectoderm-derived cells in culture, and promote wound healing in soft tissue, cartilaginous tissue and musculo-skeletal tissue. U.S. Patent No. 5,387,673 discloses biologically active fragments of FGF that retain activity.

Proteins of the TNF Family A profile derived from the TNF family is created by aligning sequences of the following TNF family members: nerve growth factor (NGF), lymphotoxin, Fas ligand, tumor necrosis factor (TNF), CD40 ligand, TRAIL, ox40 ligand, 4-1BB ligand, CD27 ligand, and CD30 ligand. The profile is designed to identify sequences of proteins that constitute new members or homologues of this family of proteins.

U.S. Patent No. 5,606,023 discloses mutant TNF proteins; U.S. Patent No. 5,597,899 and U.S. Patent No. 5,486,463 disclose TNF muteins; and U.S. Patent No. 5,652,353 discloses DNA encoding TNF- $\alpha$  muteins.

Members of the TNF family of proteins have been show in vitro to multimerize, as described in Burrows et al., Biochem. (1994) 33:12741 and Zhang et al., Mol. Cell. Biol. (1995) 154851 and bind receptors as described in Browning et al., J. Immunol. (1994) 147:1230, Androlewicz et al., J. Biol. Chem. (1992) 267:2542, and Crowe et al., Science (1994) 264:707.

In vivo, TNFs proteolytically cleave a target protein as described in Kriegel et al., Cell (1988) 53:45 and Mohler et al., Nature (1994) 370:218 and demonstrate cell proliferation and differentiation activity. T-cell or thymocyte proliferation is assayed as described in Armitage et al., Eur. J. Immunol. (1992) 22:447; Current Protocols in Immunology, ed. J.E. Coligan et al., 3.1-3.19; Takai et al., J. Immunol. (1986) 137:3494-3500, Bertagnoli et al., J. Immunol. (1990) 145:1706-1712, Bertagnoli et al., J. Immunol. (1991) 133:327-340, Bertagnoli et al., J. Immunol. (1992) 149:3778-3783, and Bowman et al., J. Immunol. (1994) 152:1756-1761. B cell proliferation and Ig secretion are assayed as described in Maliszewski, J. Immunol. (1990) 144:3028-3033, and Assays for B Cell Function: In vitro antibody production, Mond and Brunswick, Current Protocols in Immunol., Coligan Ed vol 1 pp 3.8.1-3.8.16, John Wiley and Sons, Toronto 1994, Kehrl et al., Science (1987) 238:1144 and Boussiotis et al., PNAS USA (1994) 91:7007.

Other in vivo activities include upregulation of cell surface antigens, upregulation of costimulatory molecules, and cellular aggregation/adhesion as described in Barrett et al., J. Immunol. (1991) 146:1722; Bjorck et al., Eur. J. Immunol. (1993) 23:1771; Clark et al., Annu Rev. Immunol. (1991) 9:97; Ranheim et al., J. Exp. Med. (1994) 177:925; Yellin, J. Immunol. (1994) 153:666; and Gruss et al., Blood (1994) 84:2305.

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Proliferation and differentiation of hematopoietic and lymphopoietic cells has also been shown in vivo for TNFs, using assays for embryonic differentiation and hematopoiesis as described in Johansson et al., Cellular Biology (1995) 15:141-151, Keller et al., Mol. Cell. Biol. (1993) 13:473-486, McClanahan et al., Blood (1993) 81:2903-2915 and using assays to detect stem cell survival and differentiation as described in Culture of Hematopoietic Cells, Freshney et al. eds, pp 1-21, 23-29, 139-162, 163-179, and 265-268, Wiley-Liss, Inc., New York, NY, 1994, and Hirajama et al., PNAS USA (1992) 89:5907-5911.

In vivo activities of TNFs also include lymphocyte survival and apoptosis, assayed as described in Darzynkewicz et al., Cytometry (1992) 13:795-808; Gorczca et al., Leukemia (1993) 7:659-670; Itoh et al., Cell (1991) 66:233-243; Zacharduk, J. Immunol. (1990) 145:4037-4045; Zamai et al., Cytometry (1993) 14:891-897; and Gorczyca et al., Int'l J. Oncol. (1992) 1:639-648.

Some members of the TNF family are cleaved from the cell surface; others remain membrane bound. The three-dimensional structure of TNF is discussed in Sprang and Eck, Tumor Necrosis Factors; *supra*.

TNF proteins include a transmembrane domain. The protein is cleaved into a shorter soluble version, as described in Kriegler et al., Cell (1988) 53:45-53, Perez et al., Cell (1990) 63:251-258, and Shaw et al., Cell (1986) 46:659-667. The transmembrane domain is between amino acid 46 and 77 and the cytoplasmic domain is between position 1 and 45 on the human form of TNF $\alpha$ . The 3-dimensional motifs of TNF include a sandwich of two pleated  $\beta$ -sheets. Each sheet is composed of antiparallel  $\alpha$ -strands.  $\alpha$ -Strands facing each other on opposite sites of the sandwich are connected by short polypeptide loops, as described in Van Ostade et al., Protein Engineering (1994) 7(1):5-22, and Sprang et al., Tumor Necrosis Factors; supra.

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Residues of the TNF family proteins that are involved in the  $\beta$ -sheet secondary structure have been identified as described in Van Ostade et al., Protein Engineering (1994) 7(1):5-22, and Sprang et al., Tumor Necrosis Factors; supra.

TNF receptors are disclosed in U.S. Patent No. 5,395,760. A profile derived from the TNF receptor family is created by aligning sequences of the TNF receptor family, including Apo1/Fas, TNFR I and II, death receptor3 (DR3), CD40, ox40, CD27, and CD30. Thus, the profile is designed to identify, from the nucleic acids of the invention, sequences of proteins that constitute new members or homologs of this family of proteins.

Tumor necrosis factor receptors exist in two forms in humans: p55 TNFR and p75 TNFR, both of which provide intracellular signals upon binding with a ligand. The extracellular domains of these receptor proteins are cysteine rich. The receptors can remain membrane bound, although some forms of the receptors are cleaved forming soluble receptors. The regulation, diagnostic, prognostic, and therapeutic value of soluble TNF receptors is discussed in Aderka, Cytokine and Growth Factor Reviews, (1996) 7(3):231-240.

PDGF Family U.S. Patent No. 5,326,695 discloses platelet derived growth factor agonists; bioactive portions of PDGF-B are used as agonists. U.S. Patent No. 4,845,075 discloses biologically active B-chain homodimers, and also includes variants and derivatives of the PDGF-B chain. U.S. Patent No. 5,128,321 discloses

PDGF analogs and methods of use. Proteins having the same bioactivity as PDGF are disclosed, including A and B chain proteins.

Kinase (Including MKK) Family U.S. Patent No. 5,650,501 discloses serine/threonine kinase, associated with mitotic and meiotic cell division; the protein has a kinase domain in its N-terminal and 3 PEST regions in the C-terminus. U.S. Patent No. 5,605,825 discloses human PAK65, a serine protein kinase.

The foregoing discussion provides a few examples of the protein profiles that can be compared with the nucleic acids of the invention. One skilled in the art can use these and other protein profiles to identify the genes that correlate with the nucleic acids.

# IX. Determining the Function of the Encoded Expression Products

Ribozymes, antisense constructs, dominant negative mutants, and triplex formation can be used to determine function of the expression product of an nucleic acid-related gene.

#### A. Ribozymes

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Trans-cleaving catalytic RNAs (ribozymes) are RNA molecules possessing endoribonuclease activity. Ribozymes are specifically designed for a particular target, and the target message must contain a specific nucleotide sequence. They are engineered to cleave any RNA species site-specifically in the background of cellular RNA. The cleavage event renders the mRNA unstable and prevents protein expression. Importantly, ribozymes can be used to inhibit expression of a gene of unknown function for the purpose of determining its function in an in vitro or in vivo context, by detecting the phenotypic effect.

One commonly used ribozyme motif is the hammerhead, for which the substrate sequence requirements are minimal. Design of the hammerhead ribozyme is disclosed in Usman et al., Current Opin. Struct. Biol. (1996) 6:527-533. Usman also discusses the therapeutic uses of ribozymes. Ribozymes can also be prepared and used as described in Long et al., FASEB J. (1993) 7:25; Symons, Ann. Rev. Biochem. (1992) 61:641; Perrotta et al., Biochem. (1992) 31:16-17; Ojwang et al., Proc. Natl. Acad. Sci. (USA) (1992) 89:10802-10806; and U.S. Patent No. 5,254,678. Ribozyme cleavage of HIV-I RNA is described in U.S. Patent No. 5,144,019; methods of cleaving RNA using ribozymes is described in U.S. Patent No.

5,116,742; and methods for increasing the specificity of ribozymes are described in U.S. Patent No. 5,225,337 and Koizumi et al., Nucleic Acid Res. (1989) 17:7059-7071. Preparation and use of ribozyme fragments in a hammerhead structure are also described by Koizumi et al., Nucleic Acids Res. (1989) 17:7059-7071. Preparation and use of ribozyme fragments in a hairpin structure are described by Chowrira and Burke, Nucleic Acids Res. (1992) 20:2835. Ribozymes can also be made by rolling transcription as described in Daubendiek and Kool, Nat. Biotechnol. (1997) 15(3):273-277.

The hybridizing region of the ribozyme may be modified or may be prepared as a branched structure as described in Horn and Urdea, *Nucleic Acids Res.* (1989) 17:6959-67. The basic structure of the ribozymes may also be chemically altered in ways familiar to those skilled in the art, and chemically synthesized ribozymes can be administered as synthetic oligonucleotide derivatives modified by monomeric units. In a therapeutic context, liposome mediated delivery of ribozymes improves cellular uptake, as described in Birikh *et al.*, *Eur. J. Biochem.* (1997) 245:1-16.

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Using the nucleic acid sequences of the invention and methods known in the art, ribozymes are designed to specifically bind and cut the corresponding mRNA species. Ribozymes thus provide a means to inhibit the expression of any of the proteins encoded by the disclosed nucleic acids or their full-length genes. The full-length gene need not be known in order to design and use specific inhibitory ribozymes. In the case of a nucleic acid or cDNA of unknown function, ribozymes corresponding to that nucleotide sequence can be tested in vitro for efficacy in cleaving the target transcript. Those ribozymes that effect cleavage in vitro are further tested in vivo. The ribozyme can also be used to generate an animal model for a disease, as described in Birikh et al., Eur. J. Biochem. (1997) 245:1-16. An effective ribozyme is used to determine the function of the gene of interest by blocking its transcription and detecting a change in the cell. Where the gene is found to be a mediator in a disease, an effective ribozyme is designed and delivered in a gene therapy for blocking transcription and expression of the gene.

Therapeutic and functional genomic applications of ribozymes proceed beginning with knowledge of a portion of the coding sequence of the gene to be inhibited. Thus, for many genes, a partial nucleic acid sequence provides adequate sequence for constructing an effective ribozyme. A target cleavage site is selected in

the target sequence, and a ribozyme is constructed based on the 5' and 3' nucleotide sequences that flank the cleavage site. Retroviral vectors are engineered to express monomeric and multimeric hammerhead ribozymes targeting the mRNA of the target coding sequence. These monomeric and multimeric ribozymes are tested in vitro for an ability to cleave the target mRNA. A cell line is stably transduced with the retroviral vectors expressing the ribozymes, and the transduction is confirmed by Northern blot analysis and reverse-transcription polymerase chain reaction (RT-PCR). The cells are screened for inactivation of the target mRNA by such indicators as reduction of expression of disease markers or reduction of the gene product of the target mRNA.

#### B. Antisense

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Antisense nucleic acids are designed to specifically bind to RNA, resulting in the formation of RNA-DNA or RNA-RNA hybrids, with an arrest of DNA replication, reverse transcription or messenger RNA translation. Antisense polynucleotides based on a selected nucleic acid sequence can interfere with expression of the corresponding gene. Antisense polynucleotides are typically generated within the cell by expression from antisense constructs that contain the antisense nucleic acid strand as the transcribed strand. Antisense nucleic acids will bind and/or interfere with the translation of nucleic acid-related mRNA. The expression products of control cells and cells treated with the antisense construct are compared to detect the protein product of the gene corresponding to the nucleic acid. The protein is isolated and identified using routine biochemical methods.

One rationale for using antisense methods to determine the function of the gene corresponding to a nucleic acid is the biological activity of antisense therapeutics. Antisense therapy for a variety of cancers is in clinical phase and has been discussed extensively in the literature. Reed reviewed antisense therapy directed at the Bcl-2 gene in tumors; gene transfer-mediated overexpression of Bcl-2 in tumor cell lines conferred resistance to many types of cancer drugs. (Reed, J.C., N.C.I. (1997) 89:988-990). The potential for clinical development of antisense inhibitors of ras is discussed by Cowsert, L.M., Anti-Cancer Drug Design (1997) 12:359-371. Additional important antisense targets include leukemia (Geurtz, A.M., Anti-Cancer Drug Design (1997) 12:341-358); human C-ref kinase (Monia, B.P., Anti-Cancer

Drug Design (1997) 12:327-339); and protein kinase C (McGraw et al., Anti-Cancer Drug Design (1997) 12:315-326.

Given the extensive background literature and clinical experience in antisense therapy, one skilled in the art can use selected nucleic acids of the invention as additional potential therapeutics. The choice of nucleic acid can be narrowed by first testing them for binding to "hot spot" regions of the genome of cancerous cells. If a nucleic acid is identified as binding to a "hot spot", testing the nucleic acid as an antisense compound in the corresponding cancer cells clearly is warranted.

Ogunbiyi et al., Gastroenterology (1997) 113(3):761-766 describe prognostic use of allelic loss in colon cancer; Barks et al., Genes, Chromosomes, and Cancer (1997) 19(4):278-285 describe increased chromosome copy number detected by FISH in malignant melanoma; Nishizake et al., Genes, Chromosomes, and Cancer (1997) 19(4):267-272 describe genetic alterations in primary breast cancer and their metastases and direct comparison using modified comparative genome hybridization; and Elo et al., Cancer Research (1997) 57(16):3356-3359 disclose that loss of heterozygosity at 16z24.1-q24.2 is significantly associated with metastatic and aggressive behavior of prostate cancer.

#### C. Dominant Negative Mutations

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As an alternative method for identifying function of the nucleic acid-related gene, dominant negative mutations are readily generated for corresponding proteins that are active as homomultimers. A mutant polypeptide will interact with wild-type polypeptides (made from the other allele) and form a non-functional multimer. Thus, a mutation is in a substrate-binding domain, a catalytic domain, or a cellular localization domain. Preferably, the mutant polypeptide will be overproduced. Point mutations are made that have such an effect. In addition, fusion of different polypeptides of various lengths to the terminus of a protein can yield dominant negative mutants. General strategies are available for making dominant negative mutants. See Herskowitz, *Nature* (1987) 329:219-222. Such a technique can be used for creating a loss-of-function mutation, which is useful for determining the function of a protein.

### D. Triplex Formation

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Endogenous gene expression can also be reduced by inactivating or "knocking out" the gene or its promoter using targeted homologous recombination. (E.g., see Smithies et al., 1985, Nature 317:230-234; Thomas & Capecchi, 1987, Cell 51:503-512; Thompson et al., 1989 Cell 5:313-321; each of which is incorporated by reference herein in its entirety). For example, a mutant, non-functional gene (or a completely unrelated DNA sequence) flanked by DNA homologous to the endogenous gene (either the coding regions or regulatory regions of the gene) can be used, with or without a selectable marker and/or a negative selectable marker, to transfect cells that express that gene in vivo. Insertion of the DNA construct, via targeted homologous recombination, results in inactivation of the gene.

Alternatively, endogenous gene expression can be reduced by targeting deoxyribonucleotide sequences complementary to the regulatory region of the target gene (i.e., the gene promoter and/or enhancers) to form triple helical structures that prevent transcription of the gene in target cells in the body. (See generally, Helene, C. 1991, Anticancer Drug Des., 6(6):569-84; Helene, C., et al., 1992, Ann, N.Y. Accad. Sci., 660:27-36; and Maher, L.J., 1992, Bioassays 14(12):807-15).

Nucleic acid molecules to be used in triple helix formation for the inhibition of transcription are preferably single stranded and composed of deoxyribonucleotides. The base composition of these oligonucleotides should promote triple helix formation via Hoogsteen base-pairing rules, which generally require sizable stretches of either purines or pyrimidines to be present on one strand of a duplex. Nucleotide sequences may be pyrimidine-based, which will result in TAT and CGC triplets across the three associated strands of the resulting triple helix. The pyrimidine-rich molecules provide base complementarity to a purine-rich region of a single strand of the duplex in a parallel orientation to that strand. In addition, nucleic acid molecules may be chosen that are purine-rich, for example, containing a stretch of G residues. These molecules will form a triple helix with a DNA duplex that is rich in GC pairs, in which the majority of the purine residues are located on a single strand of the targeted duplex, resulting in CGC triplets across the three strands in the triplex.

Alternatively, the potential sequences that can be targeted for triple helix formation may be increased by creating a so called "switchback" nucleic acid molecule. Switchback molecules are synthesized in an alternating 5'-3', 3'-5' manner, such that they base pair with first one strand of a duplex and then the other,

eliminating the necessity for a sizable stretch of either purines or pyrimidines to be present on one strand of a duplex.

Antisense RNA and DNA, ribozyme, and triple helix molecules of the invention may be prepared by any method known in the art for the synthesis of DNA and RNA molecules. These include techniques for chemically synthesizing oligodeoxyribonucleotides and oligoribonucleotides well known in the art such as for example solid phase phosphoramidite chemical synthesis. Alternatively, RNA molecules may be generated by *in vitro* and *in vivo* transcription of DNA sequences encoding the antisense RNA molecule. Such DNA sequences may be incorporated into a wide variety of vectors which incorporate suitable RNA polymerase promoters such as the T7 or SP6 polymerase promoters. Alternatively, antisense cDNA constructs that synthesize antisense RNA constitutively or inducibly, depending on the promoter used, can be introduced stably into cell lines.

Moreover, various well known modifications to nucleic acid molecules may be introduced as a means of increasing intracellular stability and half-life. Possible modifications include but are not limited to the addition of flanking sequences of ribonucleotides or deoxyribonucleotides to the 5' and/or 3' ends of the molecule or the use of phosphorothioate or 2' O-methyl rather than phosphodiesterase linkages within the oligodeoxyribonucleotide backbone.

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#### X. <u>Diagnostic & Prognostic Assays and Drug Screening Methods</u>

The present invention provides method for determining whether a subject is at risk for developing a disease or condition characterized by unwanted cell proliferation by detecting the disclosed biomarkers, i.e., the disclosed nucleic acid markers (SEQ ID Nos: 1-544) and/or polypeptide markers for colon cancer encoded thereby.

In clinical applications, human tissue samples can be screened for the presence and/or absence of the biomarkers identified herein. Such samples could consist of needle biopsy cores, surgical resection samples, lymph node tissue, or serum. For example, these methods include obtaining a biopsy, which is optionally fractionated by cryostat sectioning to enrich tumor cells to about 80% of the total cell population. In certain embodiments, nucleic acids extracted from these samples may be amplified using techniques well known in the art. The levels of selected markers detected

would be compared with statistically valid groups of metastatic, non-metastatic malignant, benign, or normal colon tissue samples.

In one embodiment, the diagnostic method comprises determining whether a subject has an abnormal mRNA and/or protein level of the disclosed markers, such as by Northern blot analysis, reverse transcription-polymerase chain reaction (RT-PCR), in situ hybridization, immunoprecipitation, Western blot hybridization, or immunohistochemistry. According to the method, cells are obtained from a subject and the levels of the disclosed biomarkers, protein or mRNA level, is determined and compared to the level of these markers in a healthy subject. An abnormal level of the biomarker polypeptide or mRNA levels is likely to be indicative of cancer such as colon cancer.

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Accordingly, in one aspect, the invention provides probes and primers that are specific to the unique nucleic acid markers disclosed herein. Accordingly, the nucleic acid probes comprise a nucleotide sequence at least 12 nucleotides in length, preferably at least 15 nucleotides, more preferably, 25 nucleotides, and most preferably at least 40 nucleotides, and up to all or nearly all of the coding sequence which is complementary to a portion of the coding sequence of a marker nucleic acid sequence, which nucleic acid sequence is represented by SEQ ID Nos: 1-544 or a sequence complementary thereto.

In one embodiment, the method comprises using a nucleic acid probe to determine the presence of cancerous cells in a tissue from a patient. Specifically, the method comprises:

1. providing a nucleic acid probe comprising a nucleotide sequence at least 12 nucleotides in length, preferably at least 15 nucleotides, more preferably, 25 nucleotides, and most preferably at least 40 nucleotides, and up to all or nearly all of the coding sequence which is complementary to a portion of the coding sequence of a nucleic acid sequence represented by SEQ ID Nos: 1-544 or a sequence complementary thereto and is differentially expressed in tumors cells, such as colon cancer cells;

 obtaining a tissue sample from a patient potentially comprising cancerous cells;

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 providing a second tissue sample containing cells substantially all of which are non-cancerous;

contacting the nucleic acid probe under stringent conditions with RNA of each of said first and second tissue samples (e.g., in a Northern blot or in situ hybridization assay); and comparing (a) the amount of hybridization of the probe with RNA of the first tissue sample, with (b) the amount of hybridization of the probe with RNA of the second tissue sample; wherein a statistically significant difference in the

amount of hybridization with the RNA of the first tissue sample as compared to the amount of hybridization with the RNA of the second tissue sample is indicative of the presence of cancerous cells in the first tissue sample.

In one aspect, the method comprises in situ hybridization with a probe derived from a given marker nucleic acid sequence, which nucleic acid sequence is represented by SEQ ID Nos: 1-544 or a sequence complementary thereto. The method comprises contacting the labeled hybridization probe with a sample of a given type of tissue potentially containing cancerous or pre-cancerous cells as well as normal cells, and determining whether the probe labels some cells of the given tissue type to a degree significantly different (e.g., by at least a factor of two, or at least a factor of five, or at least a factor of twenty, or at least a factor of fifty) than the degree to which it labels other cells of the same tissue type.

Also within the invention is a method of determining the phenotype of a test cell from a given human tissue, e.g., whether the cell is (a) normal, or (b) cancerous or precancerous, by contacting the mRNA of a test cell with a nucleic acid probe at least 12 nucleotides in length, preferably at least 15 nucleotides, more preferably at least 25 nucleotides, and most preferably at least 40 nucleotides, and up to all or nearly all of a sequence which is complementary to a portion of the coding sequence of a nucleic acid sequence represented by SEQ ID Nos: 1-544 or a sequence complementary thereto, and which is differentially expressed in tumor cells as compared to normal cells of the given tissue type; and determining the approximate amount of hybridization of the probe to the mRNA, an amount of hybridization either more or less than that seen with the mRNA of a normal cell of that tissue type being indicative that the test cell is cancerous or pre-cancerous.

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Alternatively, the above diagnostic assays may be carried out using antibodies to detect the protein product encoded by the marker nucleic acid sequence, which nucleic acid sequence is represented by SEQ ID Nos: 1-544 or a sequence complementary thereto. Accordingly, in one embodiment, the assay would include contacting the proteins of the test cell with an antibody specific for the gene product of a nucleic acid represented by SEQ ID Nos: 1-544 or a sequence complementary thereto, the marker nucleic acid being one which is expressed at a given control level in normal cells of the same tissue type as the test cell, and determining the approximate amount of immunocomplex formation by the antibody and the proteins of the test cell, wherein a statistically significant difference in the amount of the immunocomplex formed with the proteins of a test cell as compared to a normal cell of the same tissue type is an indication that the test cell is cancerous or pre-cancerous.

Another such method includes the steps of: providing an antibody specific for the gene product of a marker nucleic acid sequence represented by SEQ ID Nos 1-544, the gene product being present in cancerous tissue of a given tissue type (e.g., colon tissue) at a level more or less than the level of the gene product in noncancerous tissue of the same tissue type; obtaining from a patient a first sample of tissue of the given tissue type, which sample potentially includes cancerous cells; providing a second sample of tissue of the same tissue type (which may be from the same patient or from a normal control, e.g. another individual or cultured cells), this second sample containing normal cells and essentially no cancerous cells; contacting the antibody with protein (which may be partially purified, in lysed but unfractionated cells, or in situ) of the first and second samples under conditions permitting immunocomplex formation between the antibody and the marker nucleic acid sequence product present in the samples; and comparing (a) the amount of immunocomplex formation in the first sample, with (b) the amount of immunocomplex formation in the second sample, wherein a statistically significant difference in the amount of immunocomplex formation in the first sample less as compared to the amount of immunocomplex formation in the second sample is indicative of the presence of cancerous cells in the first sample of tissue.

The subject invention further provides a method of determining whether a cell sample obtained from a subject possesses an abnormal amount of marker polypeptide which comprises (a) obtaining a cell sample from the subject, (b) quantitatively

determining the amount of the marker polypeptide in the sample so obtained, and (c) comparing the amount of the marker polypeptide so determined with a known standard, so as to thereby determine whether the cell sample obtained from the subject possesses an abnormal amount of the marker polypeptide. Such marker polypeptides may be detected by immunohistochemical assays, dot-blot assays, ELISA and the like.

Immunoassays are commonly used to quantitate the levels of proteins in cell samples, and many other immunoassay techniques are known in the art. The invention is not limited to a particular assay procedure, and therefore is intended to include both homogeneous and heterogeneous procedures. Exemplary immunoassays which can be conducted according to the invention include fluorescence polarization immunoassay (FPIA), fluorescence immunoassay (FIA), enzyme immunoassay (EIA), nephelometric inhibition immunoassay (NIA), enzyme linked immunosorbent assay (ELISA), and radioimmunoassay (RIA). An indicator moiety, or label group, can be attached to the subject antibodies and is selected so as to meet the needs of various uses of the method which are often dictated by the availability of assay equipment and compatible immunoassay procedures. General techniques to be used in performing the various immunoassays noted above are known to those of ordinary skill in the art.

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In another embodiment, the level of the encoded product, i.e., the product encoded by SEQ ID Nos 1-544 or a sequence complementary thereto, in a biological fluid (e.g., blood or urine) of a patient may be determined as a way of monitoring the level of expression of the marker nucleic acid sequence in cells of that patient. Such a method would include the steps of obtaining a sample of a biological fluid from the patient, contacting the sample (or proteins from the sample) with an antibody specific for a encoded marker polypeptide, and determining the amount of immune complex formation by the antibody, with the amount of immune complex formation being indicative of the level of the marker encoded product in the sample. This determination is particularly instructive when compared to the amount of immune complex formation by the same antibody in a control sample taken from a normal individual or in one or more samples previously or subsequently obtained from the same person.

In another embodiment, the method can be used to determine the amount of marker polypeptide present in a cell, which in turn can be correlated with progression

of a hyperproliferative disorder, e.g., colon cancer. The level of the marker polypeptide can be used predictively to evaluate whether a sample of cells contains cells which are, or are predisposed towards becoming, transformed cells. Moreover, the subject method can be used to assess the phenotype of cells which are known to be transformed, the phenotyping results being useful in planning a particular therapeutic regimen. For instance, very high levels of the marker polypeptide in sample cells is a powerful diagnostic and prognostic marker for a cancer, such as colon cancer. The observation of marker polypeptide level can be utilized in decisions regarding, e.g., the use of more aggressive therapies.

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As set out above, one aspect of the present invention relates to diagnostic assays for determining, in the context of cells isolated from a patient, if the level of a marker polypeptide is significantly reduced in the sample cells. The term "significantly reduced" refers to a cell phenotype wherein the cell possesses a reduced cellular amount of the marker polypeptide relative to a normal cell of similar tissue origin. For example, a cell may have less than about 50%, 25%, 10%, or 5% of the marker polypeptide that a normal control cell. In particular, the assay evaluates the level of marker polypeptide in the test cells, and, preferably, compares the measured level with marker polypeptide detected in at least one control cell, e.g., a normal cell and/or a transformed cell of known phenotype.

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Of particular importance to the subject invention is the ability to quantitate the level of marker polypeptide as determined by the number of cells associated with a normal or abnormal marker polypeptide level. The number of cells with a particular marker polypeptide phenotype may then be correlated with patient prognosis. In one embodiment of the invention, the marker polypeptide phenotype of the lesion is determined as a percentage of cells in a biopsy which are found to have abnormally high/low levels of the marker polypeptide. Such expression may be detected by immunohistochemical assays, dot-blot assays, ELISA and the like.

Where tissue samples are employed, immunohistochemical staining may be used to determine the number of cells having the marker polypeptide phenotype. For such staining, a multiblock of tissue is taken from the biopsy or other tissue sample and subjected to proteolytic hydrolysis, employing such agents as protease K or pepsin. In certain embodiments, it may be desirable to isolate a nuclear fraction from the sample cells and detect the level of the marker polypeptide in the nuclear fraction.

The tissue samples are fixed by treatment with a reagent such as formalin, glutaraldehyde, methanol, or the like. The samples are then incubated with an antibody, preferably a monoclonal antibody, with binding specificity for the marker polypeptides. This antibody may be conjugated to a label for subsequent detection of binding. Samples are incubated for a time sufficient for formation of the immunocomplexes. Binding of the antibody is then detected by virtue of a label conjugated to this antibody. Where the antibody is unlabeled, a second labeled antibody may be employed, e.g., which is specific for the isotype of the anti-marker polypeptide antibody. Examples of labels which may be employed include radionuclides, fluorescers, chemiluminescers, enzymes and the like.

Where enzymes are employed, the substrate for the enzyme may be added to the samples to provide a colored or fluorescent product. Examples of suitable enzymes for use in conjugates include horseradish peroxidase, alkaline phosphatase, malate dehydrogenase and the like. Where not commercially available, such antibody-enzyme conjugates are readily produced by techniques known to those skilled in the art.

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In one embodiment, the assay is performed as a dot blot assay. The dot blot assay finds particular application where tissue samples are employed as it allows determination of the average amount of the marker polypeptide associated with a single cell by correlating the amount of marker polypeptide in a cell-free extract produced from a predetermined number of cells.

It is well established in the cancer literature that tumor cells of the same type (e.g., breast and/or colon tumor cells) may not show uniformly increased expression of individual oncogenes or uniformly decreased expression of individual tumor suppressor genes. There may also be varying levels of expression of a given marker gene even between cells of a given type of cancer, further emphasizing the need for reliance on a battery of tests rather than a single test. Accordingly, in one aspect, the invention provides for a battery of tests utilizing a number of probes of the invention, in order to improve the reliability and/or accuracy of the diagnostic test.

In one embodiment, the present invention also provides a method wherein nucleic acid probes are immobilized on a DNA chip in an organized array.

Oligonucleotides can be bound to a solid support by a variety of processes, including lithography. For example a chip can hold up to 250,000 oligonucleotides (GeneChip,

Affymetrix). These nucleic acid probes comprise a nucleotide sequence at least about 12 nucleotides in length, preferably at least about 15 nucleotides, more preferably at least about 25 nucleotides, and most preferably at least about 40 nucleotides, and up to all or nearly all of a sequence which is complementary to a portion of the coding sequence of a marker nucleic acid sequence represented by SEQ ID Nos: 1-544 and is differentially expressed in tumor cells, such as colon cancer cells. The present invention provides significant advantages over the available tests for various cancers, such as colon cancer, because it increases the reliability of the test by providing an array of nucleic acid markers on a single chip.

The method includes obtaining a biopsy, which is optionally fractionated by cryostat sectioning to enrich tumor cells to about 80% of the total cell population. The DNA or RNA is then extracted, amplified, and analyzed with a DNA chip to determine the presence of absence of the marker nucleic acid sequences.

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In one embodiment, the nucleic acid probes are spotted onto a substrate in a two-dimensional matrix or array. Samples of nucleic acids can be labeled and then hybridized to the probes. Double-stranded nucleic acids, comprising the labeled sample nucleic acids bound to probe nucleic acids, can be detected once the unbound portion of the sample is washed away.

The probe nucleic acids can be spotted on substrates including glass, nitrocellulose, etc. The probes can be bound to the substrate by either covalent bonds or by non-specific interactions, such as hydrophobic interactions. The sample nucleic acids can be labeled using radioactive labels, fluorophores, chromophores, etc.

Techniques for constructing arrays and methods of using these arrays are described in EP No. 0 799 897; PCT No. WO 97/29212; PCT No. WO 97/27317; EP No. 0 785 280; PCT No. WO 97/02357; U.S. Pat. No. 5,593,839; U.S. Pat. No. 5,578,832; EP No. 0 728 520; U.S. Pat. No. 5,599,695; EP No. 0 721 016; U.S. Pat. No. 5,556,752; PCT No. WO 95/22058; and U.S. Pat. No. 5,631,734.

Further, arrays can be used to examine differential expression of genes and can be used to determine gene function. For example, arrays of the instant nucleic acid sequences can be used to determine if any of the nucleic acid sequences are differentially expressed between normal cells and cancer cells, for example. High expression of a particular message in a cancer cell, which is not observed in a corresponding normal cell, can indicate a cancer specific protein.

In yet another embodiment, the invention contemplates using a panel of antibodies which are generated against the marker polypeptides of this invention, which polypeptides are encoded by SEQ ID Nos: 1-544. Such a panel of antibodies may be used as a reliable diagnostic probe for colon cancer. The assay of the present invention comprises contacting a biopsy sample containing cells, e.g., colon cells, with a panel of antibodies to one or more of the encoded products to determine the presence or absence of the marker polypeptides.

The diagnostic methods of the subject invention may also be employed as follow-up to treatment, e.g., quantitation of the level of marker polypeptides may be indicative of the effectiveness of current or previously employed cancer therapies as well as the effect of these therapies upon patient prognosis.

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Accordingly, the present invention makes available diagnostic assays and reagents for detecting gain and/or loss of marker polypeptides from a cell in order to aid in the diagnosis and phenotyping of proliferative disorders arising from, for example, tumorigenic transformation of cells.

The diagnostic assays described above can be adapted to be used as prognostic assays, as well. Such an application takes advantage of the sensitivity of the assays of the invention to events which take place at characteristic stages in the progression of a tumor. For example, a given marker gene may be up- or downregulated at a very early stage, perhaps before the cell is irreversibly committed to developing into a malignancy, while another marker gene may be characteristically up or down regulated only at a much later stage. Such a method could involve the steps of contacting the mRNA of a test cell with a nucleic acid probe derived from a given marker nucleic acid which is expressed at different characteristic levels in cancerous or precancerous cells at different stages of tumor progression, and determining the approximate amount of hybridization of the probe to the mRNA of the cell, such amount being an indication of the level of expression of the gene in the cell, and thus an indication of the stage of tumor progression of the cell; alternatively, the assay can be carried out with an antibody specific for the gene product of the given marker nucleic acid, contacted with the proteins of the test cell. A battery of such tests will disclose not only the existence and location of a tumor, but also will allow the clinician to select the mode of treatment most appropriate for the tumor, and to predict the likelihood of success of that treatment.

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The methods of the invention can also be used to follow the clinical course of a tumor. For example, the assay of the invention can be applied to a tissue sample from a patient; following treatment of the patient for the cancer, another tissue sample is taken and the test repeated. Successful treatment will result in either removal of all cells which demonstrate differential expression characteristic of the cancerous or precancerous cells, or a substantial increase in expression of the gene in those cells, perhaps approaching or even surpassing normal levels.

In yet another embodiment, the invention provides methods for determining whether a subject is at risk for developing a disease, such as a predisposition to develop cancer, for example colon cancer, associated with an aberrant activity of any one of the polypeptides encoded by nucleic acids of SEQ ID Nos: 1-544, wherein the aberrant activity of the polypeptide is characterized by detecting the presence or absence of a genetic lesion characterized by at least one of (i) an alteration affecting the integrity of a gene encoding a marker polypeptides, or (ii) the mis-expression of the encoding nucleic acid. To illustrate, such genetic lesions can be detected by ascertaining the existence of at least one of (i) a deletion of one or more nucleotides from the nucleic acid sequence, (ii) an addition of one or more nucleotides to the nucleic acid sequence, (iii) a substitution of one or more nucleotides of the nucleic acid sequence, (iv) a gross chromosomal rearrangement of the nucleic acid sequence, (v) a gross alteration in the level of a messenger RNA transcript of the nucleic acid sequence, (vii) aberrant modification of the nucleic acid sequence, such as of the methylation pattern of the genomic DNA, (vii) the presence of a non-wild type splicing pattern of a messenger RNA transcript of the gene, (viii) a non-wild type level of the marker polypeptide, (ix) allelic loss of the gene, and/or (x) inappropriate post-translational modification of the marker polypeptide.

The present invention provides assay techniques for detecting lesions in the encoding nucleic acid sequence. These methods include, but are not limited to, methods involving sequence analysis, Southern blot hybridization, restriction enzyme site mapping, and methods involving detection of absence of nucleotide pairing between the nucleic acid to be analyzed and a probe.

Specific diseases or disorders, e.g., genetic diseases or disorders, are associated with specific allelic variants of polymorphic regions of certain genes, which do not necessarily encode a mutated protein. Thus, the presence of a specific

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allelic variant of a polymorphic region of a gene in a subject can render the subject susceptible to developing a specific disease or disorder. Polymorphic regions in genes, can be identified, by determining the nucleotide sequence of genes in populations of individuals. If a polymorphic region is identified, then the link with a specific disease can be determined by studying specific populations of individuals, e.g, individuals which developed a specific disease, such as colon cancer. A polymorphic region can be located in any region of a gene, e.g., exons, in coding or non coding regions of exons, introns, and promoter region.

In an exemplary embodiment, there is provided a nucleic acid composition comprising a nucleic acid probe including a region of nucleotide sequence which is capable of hybridizing to a sense or antisense sequence of a gene or naturally occurring mutants thereof, or 5' or 3' flanking sequences or intronic sequences naturally associated with the subject genes or naturally occurring mutants thereof. The nucleic acid of a cell is rendered accessible for hybridization, the probe is contacted with the nucleic acid of the sample, and the hybridization of the probe to the sample nucleic acid is detected. Such techniques can be used to detect lesions or allelic variants at either the genomic or mRNA level, including deletions, substitutions, etc., as well as to determine mRNA transcript levels.

A preferred detection method is allele specific hybridization using probes overlapping the mutation or polymorphic site and having about 5, 10, 20, 25, or 30 nucleotides around the mutation or polymorphic region. In a preferred embodiment of the invention, several probes capable of hybridizing specifically to allelic variants are attached to a solid phase support, e.g., a "chip". Mutation detection analysis using these chips comprising oligonucleotides, also termed "DNA probe arrays" is described e.g., in Cronin et al. (1996) Human Mutation 7:244. In one embodiment, a chip comprises all the allelic variants of at least one polymorphic region of a gene. The solid phase support is then contacted with a test nucleic acid and hybridization to the specific probes is detected. Accordingly, the identity of numerous allelic variants of one or more genes can be identified in a simple hybridization experiment.

In certain embodiments, detection of the lesion comprises utilizing the probe/primer in a polymerase chain reaction (PCR) (see, e.g. U.S. Patent Nos. 4,683,195 and 4,683,202), such as anchor PCR or RACE PCR, or, alternatively, in a ligase chain reaction (LCR) (see, e.g., Landegran *et al.* (1988) *Science* 241:1077-

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1080; and Nakazawa et al. (1994) PNAS 91:360-364), the latter of which can be particularly useful for detecting point mutations in the gene (see Abravaya et al. (1995) Nuc Acid Res 23:675-682). In a merely illustrative embodiment, the method includes the steps of (i) collecting a sample of cells from a patient, (ii) isolating nucleic acid (e.g., genomic, mRNA or both) from the cells of the sample, (iii) contacting the nucleic acid sample with one or more primers which specifically hybridize to a nucleic acid sequence under conditions such that hybridization and amplification of the nucleic acid (if present) occurs, and (iv) detecting the presence or absence of an amplification product, or detecting the size of the amplification product and comparing the length to a control sample. It is anticipated that PCR and/or LCR may be desirable to use as a preliminary amplification step in conjunction with any of the techniques used for detecting mutations described herein.

Alternative amplification methods include: self sustained sequence replication (Guatelli, J.C. et al., 1990, Proc. Natl. Acad. Sci. USA 87:1874-1878), transcriptional amplification system (Kwoh, D.Y. et al., 1989, Proc. Natl. Acad. Sci. USA 86:1173-1177), Q-Beta Replicase (Lizardi, P.M. et al., 1988, Bio/Technology 6:1197), or any other nucleic acid amplification method, followed by the detection of the amplified molecules using techniques well known to those of skill in the art. These detection schemes are especially useful for the detection of nucleic acid molecules if such molecules are present in very low numbers.

In a preferred embodiment of the subject assay, mutations in, or allelic variants, of a gene from a sample cell are identified by alterations in restriction enzyme cleavage patterns. For example, sample and control DNA is isolated, amplified (optionally), digested with one or more restriction endonucleases, and fragment length sizes are determined by gel electrophoresis. Moreover, the use of sequence specific ribozymes (see, for example, U.S. Patent No. 5,498,531) can be used to score for the presence of specific mutations by development or loss of a ribozyme cleavage site.

Another aspect of the invention is directed to the identification of agents capable of modulating the differentiation and proliferation of cells characterized by aberrant proliferation. In this regard, the invention provides assays for determining compounds that modulate the expression of the marker nucleic acids (SEQ ID Nos: 1-544) and/or alter for example, inhibit the bioactivity of the encoded polypeptide.

Several in vivo methods can be used to identify compounds that modulate expression of the marker nucleic acids (SEQ ID Nos: 1-544) and/or alter for example, inhibit the bioactivity of the encoded polypeptide.

Drug screening is performed by adding a test compound to a sample of cells, and monitoring the effect. A parallel sample which does not receive the test compound is also monitored as a control. The treated and untreated cells are then compared by any suitable phenotypic criteria, including but not limited to microscopic analysis, viability testing, ability to replicate, histological examination, the level of a particular RNA or polypeptide associated with the cells, the level of enzymatic activity expressed by the cells or cell lysates, and the ability of the cells to interact with other cells or compounds. Differences between treated and untreated cells indicates effects attributable to the test compound.

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Desirable effects of a test compound include an effect on any phenotype that was conferred by the cancer-associated marker nucleic acid sequence. Examples include a test compound that limits the overabundance of mRNA, limits production of the encoded protein, or limits the functional effect of the protein. The effect of the test compound would be apparent when comparing results between treated and untreated cells.

The invention thus also encompasses methods of screening for agents which inhibit expression of the nucleic acid markers (SEQ ID Nos: 1-544) in vitro, comprising exposing a cell or tissue in which the marker nucleic acid mRNA is detectable in cultured cells to an agent in order to determine whether the agent is capable of inhibiting production of the mRNA; and determining the level of mRNA in the exposed cells or tissue, wherein a decrease in the level of the mRNA after exposure of the cell line to the agent is indicative of inhibition of the marker nucleic acid mRNA production.

Alternatively, the screening method may include in vitro screening of a cell or tissue in which marker protein is detectable in cultured cells to an agent suspected of inhibiting production of the marker protein; and determining the level of the marker protein in the cells or tissue, wherein a decrease in the level of marker protein after exposure of the cells or tissue to the agent is indicative of inhibition of marker protein production.

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The invention also encompasses in vivo methods of screening for agents which inhibit expression of the marker nucleic acids, comprising exposing a mammal having tumor cells in which marker mRNA or protein is detectable to an agent suspected of inhibiting production of marker mRNA or protein; and determining the level of marker mRNA or protein in tumor cells of the exposed mammal. A decrease in the level of marker mRNA or protein after exposure of the mammal to the agent is indicative of inhibition of marker nucleic acid expression.

Accordingly, the invention provides a method comprising incubating a cell expressing the marker nucleic acids (SEQ ID Nos: 1-544) with a test compound and measuring the mRNA or protein level. The invention further provides a method for quantitatively determining the level of expression of the marker nucleic acids in a cell population, and a method for determining whether an agent is capable of increasing or decreasing the level of expression of the marker nucleic acids in a cell population. The method for determining whether an agent is capable of increasing or decreasing the level of expression of the marker nucleic acids in a cell population comprises the steps of (a) preparing cell extracts from control and agent-treated cell populations, (b) isolating the marker polypeptides from the cell extracts, (c) quantifying (e.g., in parallel) the amount of an immunocomplex formed between the marker polypeptide and an antibody specific to said polypeptide. The marker polypeptides of this invention may also be quantified by assaying for its bioactivity. Agents that induce increased the marker nucleic acid expression may be identified by their ability to increase the amount of immunocomplex formed in the treated cell as compared with the amount of the immunocomplex formed in the control cell. In a similar manner, agents that decrease expression of the marker nucleic acid may be identified by their ability to decrease the amount of the immunocomplex formed in the treated cell extract as compared to the control cell.

mRNA levels can be determined by Northern blot hybridization. mRNA levels can also be determined by methods involving PCR. Other sensitive methods for measuring mRNA, which can be used in high throughput assays, e.g., a method using a DELFIA endpoint detection and quantification method, are described, e.g., in Webb and Hurskainen (1996) Journal of Biomolecular Screening 1:119. Marker protein levels can be determined by immunoprecipitations or immunohistochemistry using an

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antibody that specifically recognizes the protein product encoded by SEQ ID Nos: 1-544.

Agents that are identified as active in the drug screening assay are candidates to be tested for their capacity to block cell proliferation activity. These agents would be useful for treating a disorder involving aberrant growth of cells, especially colon cells.

A variety of assay formats will suffice and, in light of the present disclosure, those not expressly described herein will nevertheless be comprehended by one of ordinary skill in the art. For instance, the assay can be generated in many different formats, and include assays based on cell-free systems, e.g., purified proteins or cell lysates, as well as cell-based assays which utilize intact cells.

In many drug screening programs which test libraries of compounds and natural extracts, high throughput assays are desirable in order to maximize the number of compounds surveyed in a given period of time. Assays of the present invention which are performed in cell-free systems, such as may be derived with purified or semi-purified proteins or with lysates, are often preferred as "primary" screens in that they can be generated to permit rapid development and relatively easy detection of an alteration in a molecular target which is mediated by a test compound. Moreover, the effects of cellular toxicity and/or bioavailability of the test compound can be generally ignored in the *in vitro* system, the assay instead being focused primarily on the effect of the drug on the molecular target as may be manifest in an alteration of binding affinity with other proteins or changes in enzymatic properties of the molecular target.

# A. <u>Use of Nucleic Acids as Probes in Mapping and in Tissue Profiling</u> Probes 25

Polynucleotide probes as described above, e.g., comprising at least 12 contiguous nucleotides selected from the nucleotide sequence of an nucleic acid as shown in SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, are used for a variety of purposes, including identification of human chromosomes and determining transcription levels. Additional disclosure about preferred regions of the nucleic acid sequences is found in the accompanying tables.

The nucleotide probes are labeled, for example, with a radioactive, fluorescent, biotinylated, or chemiluminescent label, and detected by well known methods appropriate for the particular label selected. Protocols for hybridizing nucleotide probes to preparations of metaphase chromosomes are also well known in the art. A nucleotide probe will hybridize specifically to nucleotide sequences in the chromosome preparations which are complementary to the nucleotide sequence of the probe. A probe that hybridizes specifically to a nucleic acid should provide a detection signal at least 5-, 10-, or 20-fold higher than the background hybridization provided with other unrelated sequences.

In a non-limiting example, commercial programs are available for identifying regions of chromosomes commonly associated with disease, such as cancer. Nucleic acids of the invention can be used to probe these regions. For example, if, through profile searching, a nucleic acid is identified as corresponding to a gene encoding a kinase, its ability to bind to a cancer-related chromosomal region will suggest its role as a kinase in one or more stages of tumor cell development/growth. Although some experimentation would be required to elucidate the role, the nucleic acid constitutes a new material for isolating a specific protein that has potential for developing a cancer diagnostic or therapeutic.

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Nucleotide probes are used to detect expression of a gene corresponding to the nucleic acid. For example, in Northern blots, mRNA is separated electrophoretically and contacted with a probe. A probe is detected as hybridizing to an mRNA species of a particular size. The amount of hybridization is quantitated to determine relative amounts of expression, for example under a particular condition. Probes are also used to detect products of amplification by polymerase chain reaction. The products of the reaction are hybridized to the probe and hybrids are detected. Probes are used for in situ hybridization to cells to detect expression. Probes can also be used in vivo for diagnostic detection of hybridizing sequences. Probes are typically labeled with a radioactive isotope. Other types of detectable labels may be used such as chromophores, fluorophores, and enzymes.

Expression of specific mRNA can vary in different cell types and can be tissue specific. This variation of mRNA levels in different cell types can be exploited with nucleic acid probe assays to determine tissue types. For example, PCR, branched DNA probe assays, or blotting techniques utilizing nucleic acid probes substantially

identical or complementary to nucleic acids of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, can determine the presence or absence of target cDNA or mRNA.

Examples of a nucleotide hybridization assay are described in Urdea et al., PCT WO92/02526 and Urdea et al., U.S. Patent No. 5,124,246, both incorporated herein by reference. The references describe an example of a sandwich nucleotide hybridization assay.

Alternatively, the Polymerase Chain Reaction (PCR) is another means for detecting small amounts of target nucleic acids, as described in Mullis *et al.*, *Meth. Enzymol.* (1987) 155:335-350; U.S. Patent No. 4,683,195; and U.S. Patent No. 4,683,202, all incorporated herein by reference. Two primer polynucleotides nucleotides hybridize with the target nucleic acids and are used to prime the reaction. The primers may be composed of sequence within or 3' and 5' to the polynucleotides of the Sequence Listing. Alternatively, if the primers are 3' and 5' to these polynucleotides, they need not hybridize to them or the complements. A thermostable polymerase creates copies of target nucleic acids from the primers using the original target nucleic acids as a template. After a large amount of target nucleic acids is generated by the polymerase, it is detected by methods such as Southern blots. When using the Southern blot method, the labeled probe will hybridize to a polynucleotide of the Sequence Listing or complement.

Furthermore, mRNA or cDNA can be detected by traditional blotting techniques described in Sambrook et al., "Molecular Cloning: A Laboratory Manual" (New York, Cold Spring Harbor Laboratory, 1989). mRNA or cDNA generated from mRNA using a polymerase enzyme can be purified and separated using gel electrophoresis. The nucleic acids on the gel are then blotted onto a solid support, such as nitrocellulose. The solid support is exposed to a labeled probe and then washed to remove any unhybridized probe. Next, the duplexes containing the labeled probe are detected. Typically, the probe is labeled with radioactivity.

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#### **Mapping**

Nucleic acids of the present invention are used to identify a chromosome on which the corresponding gene resides. Using fluorescence in situ hybridization

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(FISH) on normal metaphase spreads, comparative genomic hybridization allows total genome assessment of changes in relative copy number of DNA sequences. See Schwartz and Samad, Current Opinions in Biotechnology (1994) 8:70-74; Kallioniemi et al., Seminars in Cancer Biology (1993) 4:41-46; Valdes and Tagle, Methods in Molecular Biology (1997) 68:1, Boultwood, ed., Human Press, Totowa, NJ.

Preparations of human metaphase chromosomes are prepared using standard cytogenetic techniques from human primary tissues or cell lines. Nucleotide probes comprising at least 12 contiguous nucleotides selected from the nucleotide sequence of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, are used to identify the corresponding chromosome. The nucleotide probes are labeled, for example, with a radioactive, fluorescent, biotinylated, or chemiluminescent label, and detected by well known methods appropriate for the particular label selected. Protocols for hybridizing nucleotide probes to preparations of metaphase chromosomes are also well known in the art. A nucleotide probe will hybridize specifically to nucleotide sequences in the chromosome preparations that are complementary to the nucleotide sequence of the probe. A probe that hybridizes specifically to a target gene provides a detection signal at least 5-, 10-, or 20-fold higher than the background hybridization provided with unrelated coding sequences.

Nucleic acids are mapped to particular chromosomes using, for example, radiation hybrids or chromosome-specific hybrid panels. See Leach et al., Advances in Genetics, (1995) 33:63-99; Walter et al., Nature Genetics (1994) 7:22-28; Walter and Goodfellow, Trends in Genetics (1992) 9:352. Panels for radiation hybrid mapping are available from Research Genentics, Inc., Huntsville, Alabama, USA. Databases for markers using various panels are available via the world wide web at http://F/shgc-www.stanford.edu; and other locations. The statistical program RHMAP can be used to construct a map based on the data from radiation hybridization with a measure of the relative likelihood of one order versus another. RHMAP is available via the world wide web at http://www.sph.umich.edu/group/statgen/software.

Such mapping can be useful in identifying the function of the target gene by its proximity to other genes with known function. Function can also be assigned to the target gene when particular syndromes or diseases map to the same chromosome.

#### Tissue Profiling

The nucleic acids of the present invention can be used to determine the tissue type from which a given sample is derived. For example, a metastatic lesion is identified by its developmental organ or tissue source by identifying the expression of a particular marker of that organ or tissue. If a nucleic acid is expressed only in a specific tissue type, and a metastatic lesion is found to express that nucleic acid, then the developmental source of the lesion has been identified. Expression of a particular nucleic acid is assayed by detection of either the corresponding mRNA or the protein product. Immunological methods, such as antibody staining, are used to detect a particular protein product. Hybridization methods may be used to detect particular mRNA species, including but not limited to in situ hybridization and Northern blotting.

#### Use of Polymorphisms

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A nucleic acid will be useful in forensics, genetic analysis, mapping, and diagnostic applications if the corresponding region of a gene is polymorphic in the human population. A particular polymorphic form of the nucleic acid may be used to either identify a sample as deriving from a suspect or rule out the possibility that the sample derives from the suspect. Any means for detecting a polymorphism in a gene are used, including but not limited to electrophoresis of protein polymorphic variants, differential sensitivity to restriction enzyme cleavage, and hybridization to an allelespecific probe.

#### B. <u>Use of Nucleic Acids and Encoded Polypeptides to Raise Antibodies</u>

Expression products of a nucleic acid, the corresponding mRNA or cDNA, or the corresponding complete gene are prepared and used for raising antibodies for experimental, diagnostic, and therapeutic purposes. For nucleic acids to which a corresponding gene has not been assigned, this provides an additional method of identifying the corresponding gene. The nucleic acid or related cDNA is expressed as described above, and antibodies are prepared. These antibodies are specific to an epitope on the encoded polypeptide, and can precipitate or bind to the corresponding native protein in a cell or tissue preparation or in a cell-free extract of an in vitro expression system.

Immunogens for raising antibodies are prepared by mixing the polypeptides encoded by the nucleic acids of the present invention with adjuvants. Alternatively, polypeptides are made as fusion proteins to larger immunogenic proteins. Polypeptides are also covalently linked to other larger immunogenic proteins, such as keyhole limpet hemocyanin. Immunogens are typically administered intradermally, subcutaneously, or intramuscularly. Immunogens are administered to experimental animals such as rabbits, sheep, and mice, to generate antibodies. Optionally, the animal spleen cells are isolated and fused with myeloma cells to form hybridomas which secrete monoclonal antibodies. Such methods are well known in the art. According to another method known in the art, the nucleic acid is administered directly, such as by intramuscular injection, and expressed in vivo. The expressed protein generates a variety of protein-specific immune responses, including production of antibodies, comparable to administration of the protein.

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Preparations of polyclonal and monoclonal antibodies specific for nucleic acid-encoded proteins and polypeptides are made using standard methods known in the art. The antibodies specifically bind to epitopes present in the polypeptides encoded by a nucleic acid of SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto. In another embodiment, the antibodies specifically bind to epitopes present in a polypeptide encoded by SEQ ID Nos. 1-544. Typically, at least about 6, 8, 10, or 12 contiguous amino acids are required to form an epitope. However, epitopes which involve noncontiguous amino acids may require more, for example, at least about 15, 25, or 50 amino acids. A short sequence of a nucleic acid may then be unsuitable for use as an epitope to raise antibodies for identifying the corresponding novel protein, because of the potential for cross-reactivity with a known protein. However, the antibodies may be useful for other purposes, particularly if they identify common structural features of a known protein and a novel polypeptide encoded by a nucleic acid of the invention.

Antibodies that specifically bind to human nucleic acid-encoded polypeptides should provide a detection signal at least about 5-, 10-, or 20-fold higher than a detection signal provided with other proteins when used in Western blots or other immunochemical assays. Preferably, antibodies that specifically bind nucleic acid T-

encoded polypeptides do not detect other proteins in immunochemical assays and can immunoprecipitate nucleic acid-encoded proteins from solution.

To test for the presence of serum antibodies to the nucleic acid-encoded polypeptide in a human population, human antibodies are purified by methods well known in the art. Preferably, the antibodies are affinity purified by passing antiserum over a column to which a nucleic acid-encoded protein, polypeptide, or fusion protein is bound. The bound antibodies can then be eluted from the column, for example using a buffer with a high salt concentration.

In addition to the antibodies discussed above, genetically engineered antibody derivatives are made, such as single chain antibodies.

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Antibodies may be made by using standard protocols known in the art (See, for example, Antibodies: A Laboratory Manual ed. by Harlow and Lane (Cold Spring Harbor Press: 1988)). A mammal, such as a mouse, hamster, or rabbit can be immunized with an immunogenic form of the peptide (e.g., a mammalian polypeptide or an antigenic fragment which is capable of eliciting an antibody response, or a fusion protein as described above).

In one aspect, this invention includes monoclonal antibodies that show a subject polypeptide is highly expressed in colorectal tissue or tumor tissue, especially colon cancer tissue or colon cancer-derived cell lines. Therefore, in one embodiment, this invention provides a diagnostic tool for the analysis of expression of a subject polypeptide in general, and in particular, as a diagnostic for colon cancer.

Techniques for conferring immunogenicity on a protein or peptide include conjugation to carriers or other techniques well known in the art. An immunogenic portion of a protein can be administered in the presence of adjuvant. The progress of immunization can be monitored by detection of antibody titers in plasma or serum. Standard ELISA or other immunoassays can be used with the immunogen as antigen to assess the levels of antibodies. In a preferred embodiment, the subject antibodies are immunospecific for antigenic determinants of a protein of a mammal, e.g., antigenic determinants of a protein encoded by one of SEQ ID Nos. 1-544 or closely related homologs (e.g., at least 90% identical, and more preferably at least 95% identical).

Following immunization of an animal with an antigenic preparation of a polypeptide, antisera can be obtained and, if desired, polyclonal antibodies isolated

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from the serum. To produce monoclonal antibodies, antibody-producing cells (lymphocytes) can be harvested from an immunized animal and fused by standard somatic cell fusion procedures with immortalizing cells such as myeloma cells to yield hybridoma cells. Such techniques are well known in the art, and include, for example, the hybridoma technique (originally developed by Kohler and Milstein, (1975) Nature, 256: 495-497), the human B cell hybridoma technique (Kozbar *et al.*, (1983) Immunology Today, 4: 72), and the EBV-hybridoma technique to produce human monoclonal antibodies (Cole et al., (1985) Monoclonal Antibodies and Cancer Therapy, Alan R. Liss, Inc. pp. 77-96). Hybridoma cells can be screened immunochemically for production of antibodies specifically reactive with a polypeptide of the present invention and monoclonal antibodies isolated from a culture comprising such hybridoma cells.

The term antibody as used herein is intended to include fragments thereof which are also specifically reactive with one of the subject polypeptides. Antibodies can be fragmented using conventional techniques and the fragments screened for utility in the same manner as described above for whole antibodies. For example,  $F(ab)_2$  fragments can be generated by treating antibody with pepsin. The resulting  $F(ab)_2$  fragment can be treated to reduce disulfide bridges to produce Fab fragments. The antibody of the present invention is further intended to include bispecific, single-chain, and chimeric and humanized molecules having affinity for a polypeptide conferred by at least one CDR region of the antibody. In preferred embodiments, the antibodies, the antibody further comprises a label attached thereto and able to be detected, (e.g., the label can be a radioisotope, fluorescent compound, chemiluminescent compound, enzyme, or enzyme co-factor).

Antibodies can be used, e.g., to monitor protein levels in an individual for determining, e.g., whether a subject has a disease or condition, such as colon cancer, associated with an aberrant protein level, or allowing determination of the efficacy of a given treatment regimen for an individual afflicted with such a disorder. The level of polypeptides may be measured from cells in bodily fluid, such as in blood samples.

Another application of antibodies of the present invention is in the immunological screening of cDNA libraries constructed in expression vectors such as gt11, gt18-23, ZAP, and ORF8. Messenger libraries of this type, having coding sequences inserted in the correct reading frame and orientation, can produce fusion

proteins. For instance, gtll will produce fusion proteins whose amino termini consist of ß-galactosidase amino acid sequences and whose carboxyl termini consist of a foreign polypeptide. Antigenic epitopes of a protein, e.g., other orthologs of a particular protein or other paralogs from the same species, can then be detected with antibodies, as, for example, reacting nitrocellulose filters lifted from infected plates with antibodies. Positive phage detected by this assay can then be isolated from the infected plate. Thus, the presence of homologs can be detected and cloned from other animals, as can alternate isoforms (including splicing variants) from humans.

In another embodiment, a panel of monoclonal antibodies may be used, wherein each of the epitope's involved functions are represented by a monoclonal antibody. Loss or perturbation of binding of a monoclonal antibody in the panel would be indicative of a mutational attention of the protein and thus of the corresponding gene.

#### C. Differential Expression

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The present invention also provides a method to identify abnormal or diseased tissue in a human. For nucleic acids corresponding to profiles of protein families as described above, the choice of tissue may be dictated by the putative biological function. The expression of a gene corresponding to a specific nucleic acid is compared between a first tissue that is suspected of being diseased and a second, normal tissue of the human. The normal tissue is any tissue of the human, especially those that express the target gene including, but not limited to, brain, thymus, testis, heart, prostate, placenta, spleen, small intestine, skeletal muscle, pancreas, and the mucosal lining of the colon.

The tissue suspected of being abnormal or diseased can be derived from a different tissue type of the human, but preferably it is derived from the same tissue type; for example an intestinal polyp or other abnormal growth should be compared with normal intestinal tissue. A difference between the target gene, mRNA, or protein in the two tissues which are compared, for example in molecular weight, amino acid or nucleotide sequence, or relative abundance, indicates a change in the gene, or a gene which regulates it, in the tissue of the human that was suspected of being diseased.

The target genes in the two tissues are compared by any means known in the art. For example, the two genes are sequenced, and the sequence of the gene in the tissue suspected of being diseased is compared with the gene sequence in the normal tissue. The target genes, or portions thereof, in the two tissues are amplified, for example using nucleotide primers based on the nucleotide sequence shown in the Sequence Listing, using the polymerase chain reaction. The amplified genes or portions of genes are hybridized to nucleotide probes selected from a corresponding nucleotide sequence shown SEQ ID No. 1-544. A difference in the nucleotide sequence of the target gene in the tissue suspected of being diseased compared with the normal nucleotide sequence suggests a role of the nucleic acid-encoded proteins in the disease, and provides a lead for preparing a therapeutic agent. The nucleotide probes are labeled by a variety of methods, such as radiolabeling, biotinylation, or labeling with fluorescent or chemiluminescent tags, and detected by standard methods known in the art.

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Alternatively, target mRNA in the two tissues is compared. PolyA+RNA is isolated from the two tissues as is known in the art. For example, one of skill in the art can readily determine differences in the size or amount of target mRNA transcripts between the two tissues using Northern blots and nucleotide probes selected from the nucleotide sequence shown in the Sequence Listing. Increased or decreased expression of a target mRNA in a tissue sample suspected of being diseased, compared with the expression of the same target mRNA in a normal tissue, suggests that the expressed protein has a role in the disease, and also provides a lead for preparing a therapeutic agent.

Any method for analyzing proteins is used to compare two nucleic acidencoded proteins from matched samples. The sizes of the proteins in the two tissues are compared, for example, using antibodies of the present invention to detect nucleic acid-encoded proteins in Western blots of protein extracts from the two tissues. Other changes, such as expression levels and subcellular localization, can also be detected immunologically, using antibodies to the corresponding protein. A higher or lower level of nucleic acid-encoded protein expression in a tissue suspected of being diseased, compared with the same nucleic acid-encoded protein expression level in a normal tissue, is indicative that the expressed protein has a role in the disease, and provides another lead for preparing a therapeutic agent.

Similarly, comparison of gene sequences or of gene expression products, e.g., mRNA and protein, between a human tissue that is suspected of being diseased and a normal tissue of a human, are used to follow disease progression or remission in the human. Such comparisons of genes, mRNA, or protein are made as described above.

For example, increased or decreased expression of the target gene in the tissue suspected of being neoplastic can indicate the presence of neoplastic cells in the tissue. The degree of increased expression of the target gene in the neoplastic tissue relative to expression of the gene in normal tissue, or differences in the amount of increased expression of the target gene in the neoplastic tissue over time, is used to assess the progression of the neoplasia in that tissue or to monitor the response of the neoplastic tissue to a therapeutic protocol over time.

The expression pattern of any two cell types can be compared, such as low and high metastatic tumor cell lines, or cells from tissue which have and have not been exposed to a therapeutic agent. A genetic predisposition to disease in a human is detected by comparing an target gene, mRNA, or protein in a fetal tissue with a normal target gene, mRNA, or protein. Fetal tissues that are used for this purpose include, but are not limited to, amniotic fluid, chorionic villi, blood, and the blastomere of an in vitro-fertilized embryo. The comparable normal target gene is obtained from any tissue. The mRNA or protein is obtained from a normal tissue of a human in which the target gene is expressed. Differences such as alterations in the nucleotide sequence or size of the fetal target gene or mRNA, or alterations in the molecular weight, amino acid sequence, or relative abundance of fetal target protein, can indicate a germline mutation in the target gene of the fetus, which indicates a genetic predisposition to disease.

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# D. <u>Use of Nucleic Acids, and Encoded Polypeptides to Screen for Peptide Analogs and Antagonists</u>

Polypeptides encoded by the instant nucleic acids, e.g., SEQ ID Nos. 1-544, preferably SEQ ID Nos. 1-168, even more preferably SEQ ID Nos. 1-35, or a sequence complementary thereto, and corresponding full length genes can be used to screen peptide libraries to identify binding partners, such as receptors, from among the encoded polypeptides.

A library of peptides may be synthesized following the methods disclosed in U.S. Pat. No. 5,010,175, and in PCT WO 91/17823. As described below in brief, one prepares a mixture of peptides, which is then screened to identify the peptides exhibiting the desired signal transduction and receptor binding activity. In the '175 method, a suitable peptide synthesis support (e.g., a resin) is coupled to a mixture of appropriately protected, activated amino acids. The concentration of each amino acid in the reaction mixture is balanced or adjusted in inverse proportion to its coupling reaction rate so that the product is an equimolar mixture of amino acids coupled to the starting resin. The bound amino acids are then deprotected, and reacted with another balanced amino acid mixture to form an equimolar mixture of all possible dipeptides. This process is repeated until a mixture of peptides of the desired length (e.g., hexamers) is formed. Note that one need not include all amino acids in each step; one may include only one or two amino acids in some steps (e.g., where it is known that a particular amino acid is essential in a given position), thus reducing the complexity of the mixture. After the synthesis of the peptide library is completed, the mixture of peptides is screened for binding to the selected polypeptide. The peptides are then tested for their ability to inhibit or enhance activity. Peptides exhibiting the desired activity are then isolated and sequenced.

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The method described in WO 91/17823 is similar. However, instead of reacting the synthesis resin with a mixture of activated amino acids, the resin is divided into twenty equal portions (or into a number of portions corresponding to the number of different amino acids to be added in that step), and each amino acid is coupled individually to its portion of resin. The resin portions are then combined, mixed, and again divided into a number of equal portions for reaction with the second amino acid. In this manner, each reaction may be easily driven to completion. Additionally, one may maintain separate "subpools" by treating portions in parallel, rather than combining all resins at each step. This simplifies the process of determining which peptides are responsible for any observed receptor binding or signal transduction activity.

In such cases, the subpools containing, e.g., 1-2,000 candidates each are exposed to one or more polypeptides of the invention. Each subpool that produces a positive result is then resynthesized as a group of smaller subpools (sub-subpools) containing, e.g., 20-100 candidates, and reassayed. Positive sub-subpools may be

resynthesized as individual compounds, and assayed finally to determine the peptides that exhibit a high binding constant. These peptides can be tested for their ability to inhibit or enhance the native activity. The methods described in WO 91/7823 and U.S. Patent No. 5,194,392 (herein incorporated by reference) enable the preparation of such pools and subpools by automated techniques in parallel, such that all synthesis and resynthesis may be performed in a matter of days.

Peptide agonists or antagonists are screened using any available method, such as signal transduction, antibody binding, receptor binding, mitogenic assays, chemotaxis assays, etc. The methods described herein are presently preferred. The assay conditions ideally should resemble the conditions under which the native activity is exhibited *in vivo*, that is, under physiologic pH, temperature, and ionic strength. Suitable agonists or antagonists will exhibit strong inhibition or enhancement of the native activity at concentrations that do not cause toxic side effects in the subject. Agonists or antagonists that compete for binding to the native polypeptide may require concentrations equal to or greater than the native concentration, while inhibitors capable of binding irreversibly to the polypeptide may be added in concentrations on the order of the native concentration.

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The end results of such screening and experimentation will be at least one novel polypeptide binding partner, such as a receptor, encoded by a nucleic acid of the invention, and at least one peptide agonist or antagonist of the novel binding partner. Such agonists and antagonists can be used to modulate, enhance, or inhibit receptor function in cells to which the receptor is native, or in cells that possess the receptor as a result of genetic engineering. Further, if the novel receptor shares biologically important characteristics with a known receptor, information about agonist/antagonist binding may help in developing improved agonists/antagonists of the known receptor.

# E. <u>Pharmaceutical Compositions and Therapeutic Uses</u>

Pharmaceutical compositions can comprise polypeptides, antibodies, or polynucleotides of the claimed invention. The pharmaceutical compositions will comprise a therapeutically effective amount of either polypeptides, antibodies, or polynucleotides of the claimed invention.

The term "therapeutically effective amount" as used herein refers to an amount of a therapeutic agent to treat, ameliorate, or prevent a desired disease or condition, or

to exhibit a detectable therapeutic or preventative effect. The effect can be detected by, for example, chemical markers or antigen levels. Therapeutic effects also include reduction in physical symptoms, such as decreased body temperature. The precise effective amount for a subject will depend upon the subject's size and health, the nature and extent of the condition, and the therapeutics or combination of therapeutics selected for administration. Thus, it is not useful to specify an exact effective amount in advance. However, the effective amount for a given situation can be determined by routine experimentation and is within the judgment of the clinician.

For purposes of the present invention, an effective dose will be from about 0.01 mg/kg to 50 mg/kg or 0.05 mg/kg to about 10 mg/kg of the DNA constructs in the individual to which it is administered.

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A pharmaceutical composition can also contain a pharmaceutically acceptable carrier. The term "pharmaceutically acceptable carrier" refers to a carrier for administration of a therapeutic agent, such as antibodies or a polypeptide, genes, and other therapeutic agents. The term refers to any pharmaceutical carrier that does not itself induce the production of antibodies harmful to the individual receiving the composition, and which may be administered without undue toxicity. Suitable carriers may be large, slowly metabolized macromolecules such as proteins, polysaccharides, polylactic acids, polyglycolic acids, polymeric amino acids, amino acid copolymers, and inactive virus particles. Such carriers are well known to those of ordinary skill in the art.

Pharmaceutically acceptable salts can be used therein, for example, mineral acid salts such as hydrochlorides, hydrobromides, phosphates, sulfates, and the like; and the salts of organic acids such as acetates, propionates, malonates, benzoates, and the like. A thorough discussion of pharmaceutically acceptable excipients is available in *Remington's Pharmaceutical Sciences* (Mack Pub. Co., N.J. 1991).

Pharmaceutically acceptable carriers in therapeutic compositions may contain liquids such as water, saline, glycerol and ethanol. Additionally, auxiliary substances, such as wetting or emulsifying agents, pH buffering substances, and the like, may be present in such vehicles. Typically, the therapeutic compositions are prepared as injectables, either as liquid solutions or suspensions; solid forms suitable for solution in, or suspension in, liquid vehicles prior to injection may also be prepared.

Liposomes are included within the definition of a pharmaceutically acceptable carrier.

#### Delivery Methods

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Once formulated, the nucleic acid compositions of the invention can be (1) administered directly to the subject; (2) delivered ex vivo, to cells derived from the subject; or (3) delivered in vitro for expression of recombinant proteins.

Direct delivery of the compositions will generally be accomplished by injection, either subcutaneously, intraperitoneally, intravenously or intramuscularly, or delivered to the interstitial space of a tissue. The compositions can also be administered into a tumor or lesion. Other modes of administration include oral and pulmonary administration, suppositories, and transdermal applications, needles, and gene guns or hyposprays. Dosage treatment may be a single dose schedule or a multiple dose schedule.

Methods for the ex vivo delivery and reimplantation of transformed cells into a subject are known in the art and described in e.g., International Publication No. WO 93/14778. Examples of cells useful in ex vivo applications include, for example, stem cells, particularly hematopoetic, lymph cells, macrophages, dendritic cells, or tumor cells.

Generally, delivery of nucleic acids for both ex vivo and in vitro applications can be accomplished by, for example, dextran-mediated transfection, calcium phosphate precipitation, polybrene mediated transfection, protoplast fusion, electroporation, encapsulation of the polynucleotide(s) in liposomes, and direct microinjection of the DNA into nuclei, all well known in the art.

Once a subject gene has been found to correlate with a proliferative disorder, such as neoplasia, dysplasia, and hyperplasia, the disorder may be amenable to treatment by administration of a therapeutic agent based on the nucleic acid or corresponding polypeptide.

Preparation of antisense polypeptides is discussed above. Neoplasias that are treated with the antisense composition include, but are not limited to, cervical cancers, melanomas, colorectal adenocarcinomas, Wilms' tumor, retinoblastoma, sarcomas, myosarcomas, lung carcinomas, leukemias, such as chronic myelogenous leukemia, promyelocytic leukemia, monocytic leukemia, and myeloid leukemia, and lymphomas, such as histiocytic lymphoma. Proliferative disorders that are treated with the therapeutic composition include disorders such as anhydric hereditary

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ectodermal dysplasia, congenital alveolar dysplasia, epithelial dysplasia of the cervix, fibrous dysplasia of bone, and mammary dysplasia. Hyperplasias, for example, endometrial, adrenal, breast, prostate, or thyroid hyperplasias or pseudoepitheliomatous hyperplasia of the skin, are treated with antisense therapeutic compositions. Even in disorders in which mutations in the corresponding gene are not implicated, downregulation or inhibition of nucleic acid-related gene expression can have therapeutic application. For example, decreasing nucleic acid-related gene expression can help to suppress tumors in which enhanced expression of the gene is implicated.

Both the dose of the antisense composition and the means of administration are determined based on the specific qualities of the therapeutic composition, the condition, age, and weight of the patient, the progression of the disease, and other relevant factors. Administration of the therapeutic antisense agents of the invention includes local or systemic administration, including injection, oral administration, particle gun or catheterized administration, and topical administration. Preferably, the therapeutic antisense composition contains an expression construct comprising a promoter and a polynucleotide segment of at least about 12, 22, 25, 30, or 35 contiguous nucleotides of the antisense strand of a nucleic acid. Within the expression construct, the polynucleotide segment is located downstream from the promoter, and transcription of the polynucleotide segment initiates at the promoter.

Various methods are used to administer the therapeutic composition directly to a specific site in the body. For example, a small metastatic lesion is located and the therapeutic composition injected several times in several different locations within the body of tumor. Alternatively, arteries which serve a tumor are identified, and the therapeutic composition injected into such an artery, in order to deliver the composition directly into the tumor. A tumor that has a necrotic center is aspirated and the composition injected directly into the now empty center of the tumor. The antisense composition is directly administered to the surface of the tumor, for example, by topical application of the composition. X-ray imaging is used to assist in certain of the above delivery methods.

Receptor-mediated targeted delivery of therapeutic compositions containing an antisense polynucleotide, subgenomic polynucleotides, or antibodies to specific tissues is also used. Receptor-mediated DNA delivery techniques are described in, for

example, Findeis et al., Trends in Biotechnol. (1993) 11:202-205; Chiou et al., (1994) Gene Therapeutics: Methods And Applications Of Direct Gene Transfer (J.A. Wolff, ed.); Wu & Wu, J. Biol. Chem. (1988) 263:621-24; Wu et al., J. Biol. Chem. (1994) 269:542-46; Zenke et al., Proc. Natl. Acad. Sci. (USA) (1990) 87:3655-59; Wu et al., J. Biol. Chem. (1991) 266:338-42. Preferably, receptor-mediated targeted delivery of therapeutic compositions containing antibodies of the invention is used to deliver the antibodies to specific tissue.

Therapeutic compositions containing antisense subgenomic polynucleotides are administered in a range of about 100 ng to about 200 mg of DNA for local administration in a gene therapy protocol. Concentration ranges of about 500 ng to about 50 mg, about 1 mg to about 2 mg, about 5 mg to about 500 mg, and about 20 mg to about 100 mg of DNA can also be used during a gene therapy protocol. Factors such as method of action and efficacy of transformation and expression are considerations which will affect the dosage required for ultimate efficacy of the antisense subgenomic nucleic acids. Where greater expression is desired over a larger area of tissue, larger amounts of antisense subgenomic nucleic acids or the same amounts readministered in a successive protocol of administrations, or several administrations to different adjacent or close tissue portions of, for example, a tumor site, may be required to effect a positive therapeutic outcome. In all cases, routine experimentation in clinical trials will determine specific ranges for optimal therapeutic effect. A more complete description of gene therapy vectors, especially retroviral vectors, is contained in U.S. Serial No. 08/869,309, which is expressly incorporated herein, and in section F below.

For genes encoding polypeptides or proteins with anti-inflammatory activity, suitable use, doses, and administration are described in U.S. Patent No. 5,654,173, incorporated herein by reference. Therapeutic agents also include antibodies to proteins and polypeptides encoded by the subject nucleic acids, as described in U.S. Patent No. 5,654,173.

#### F. Gene Therapy

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The therapeutic nucleic acids of the present invention may be utilized in gene delivery vehicles. The gene delivery vehicle may be of viral or non-viral origin (see generally, Jolly, Cancer Gene Therapy (1994) 1:51-64; Kimura, Human Gene

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Therpay (1994) 5:845-852; Connelly, Human Gene Therapy (1995) 1:185-193; and Kaplitt, Nature Genetics (1994) 6:148-153). Gene therapy vehicles for delivery of constructs including a coding sequence of a therapeutic of the invention can be administered either locally or systemically. These constructs can utilize viral or non-viral vector approaches. Expression of such coding sequences can be induced using endogenous mammalian or heterologous promoters. Expression of the coding sequence can be either constitutive or regulated.

The present invention can employ recombinant retroviruses which are constructed to carry or express a selected nucleic acid molecule of interest. Retrovirus vectors that can be employed include those described in EP 0 415 731; WO 90/07936; WO 94/03622; WO 93/25698; WO 93/25234; U.S. Patent No. 5, 219,740; WO 93/11230; WO 93/10218; Vile and Hart, Cancer Res. (1993) 53:3860-3864; Vile and Hart, Cancer Res. (1993) 53:962-967; Ram et al., Cancer Res. (1993) 53:83-88; Takamiya et al., J. Neurosci. Res. (1992) 33:493-503; Baba et al., J. Neurosurg. (1993) 79:729-735; U.S. Patent no. 4,777,127; GB Patent No. 2,200,651; and EP 0 345 242. Preferred recombinant retroviruses include those described in WO 91/02805.

Packaging cell lines suitable for use with the above-described retroviral vector constructs may be readily prepared (see PCT publications WO 95/30763 and WO 92/05266), and used to create producer cell lines (also termed vector cell lines) for the production of recombinant vector particles. Within particularly preferred embodiments of the invention, packaging cell lines are made from human (such as HT1080 cells) or mink parent cell lines, thereby allowing production of recombinant retroviruses that can survive inactivation in human serum.

The present invention also employs alphavirus-based vectors that can function as gene delivery vehicles. Such vectors can be constructed from a wide variety of alphaviruses, including, for example, Sindbis virus vectors, Semliki forest virus (ATCC VR-67; ATCC VR-1247), Ross River virus (ATCC VR-373; ATCC VR-1246) and Venezuelan equine encephalitis virus (ATCC VR-923; ATCC VR-1250; ATCC VR 1249; ATCC VR-532). Representative examples of such vector systems include those described in U.S. Patent Nos. 5,091,309; 5,217,879; and 5,185,440; and PCT Publication Nos. WO 92/10578; WO 94/21792; WO 95/27069; WO 95/27044; and WO 95/07994.

Gene delivery vehicles of the present invention can also employ parvovirus such as adeno-associated virus (AAV) vectors. Representative examples include the AAV vectors disclosed by Srivastava in WO 93/09239, Samulski et al., *J. Vir.* (1989) 63:3822-3828; Mendelson et al., *Virol.* (1988) 166:154-165; and Flotte et al., *PNAS* (1993) 90:10613-10617.

Representative examples of adenoviral vectors include those described by Berkner, Biotechniques (1988) 6:616-627; Rosenfeld et al., Science (1991) 252:431-434; WO 93/19191; Kolls et al., PNAS (1994) 91:215-219; Kass-Eisler et al., PNAS (1993) 90:11498-11502; Guzman et al., Circulation (1993) 88:2838-2848; Guzman et al., Cir. Res. (1993) 73:1202-1207; Zabner et al., Cell (1993) 75:207-216; Li et al., Hum. Gene Ther. (1993) 4:403-409; Cailaud et al., Eur. J. Neurosci. (1993) 5:1287-1291; Vincent et al., Nat. Genet. (1993) 5:130-134; Jaffe et al., Nat. Genet. (1992) 1:372-378; and Levrero et al., Gene (1991) 101:195-202. Exemplary adenoviral gene therapy vectors employable in this invention also include those described in WO 94/12649, WO 93/03769; WO 93/19191; WO 94/28938; WO 95/11984 and WO 95/00655. Administration of DNA linked to killed adenovirus as described in Curiel, Hum. Gene Ther. (1992) 3:147-154 may be employed.

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Other gene delivery vehicles and methods may be employed, including polycationic condensed DNA linked or unlinked to killed adenovirus alone, for example Curiel, *Hum. Gene Ther.* (1992) 3:147-154; ligand linked DNA, for example see Wu, *J. Biol. Chem.* (1989) 264:16985-16987; eukaryotic cell delivery vehicles cells, for example see U.S. Serial No. 08/240,030, filed May 9, 1994, and U.S. Serial No. 08/404,796; deposition of photopolymerized hydrogel materials; hand-held gene transfer particle gun, as described in U.S. Patent No. 5,149,655; ionizing radiation as described in U.S. Patent No. 5,206,152 and in WO92/11033; nucleic charge neutralization or fusion with cell membranes. Additional approaches are described in Philip, *Mol. Cell Biol.* (1994) 14:2411-2418, and in Woffendin, *Proc. Natl. Acad. Sci.* (1994) 91:1581-1585.

Naked DNA may also be employed. Exemplary naked DNA introduction methods are described in WO 90/11092 and U.S. Patent No. 5,580,859. Uptake efficiency may be improved using biodegradable latex beads. DNA coated latex beads are efficiently transported into cells after endocytosis initiation by the beads. The method may be improved further by treatment of the beads to increase

hydrophobicity and thereby facilitate disruption of the endosome and release of the DNA into the cytoplasm. Liposomes that can act as gene delivery vehicles are described in U.S. Patent No. 5,422,120, PCT Nos. WO 95/13796, WO 94/23697, and WO 91/14445, and EP No. 0 524 968.

Further non-viral delivery suitable for use includes mechanical delivery systems such as the approach described in Woffendin et al., Proc. Natl. Acad. Sci. USA (1994) 91(24):11581-11585. Moreover, the coding sequence and the product of expression of such can be delivered through deposition of photopolymerized hydrogel materials. Other conventional methods for gene delivery that can be used for delivery of the coding sequence include, for example, use of hand-held gene transfer particle gun, as described in U.S. Patent No. 5,149,655; use of ionizing radiation for activating transferred gene, as described in U.S. Patent No. 5,206,152 and PCT No. WO 92/11033.

#### G. Transgenic Animals

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One aspect of the present invention relates to transgenic non-human animals having germline and/or somatic cells in which the biological activity of one or more genes are altered by a chromosomally incorporated transgene.

In a preferred embodiments, the transgene encodes a mutant protein, such as dominant negative protein which antagonizes at least a portion of the biological function of a wild-type protein.

Yet another preferred transgenic animal includes a transgene encoding an antisense transcript which, when transcribed from the transgene, hybridizes with a gene or a mRNA transcript thereof, and inhibits expression of the gene.

In one embodiment, the present invention provides a desired non-human animal or an animal (including human) cell which contains a predefined, specific and desired alteration rendering the non-human animal or animal cell predisposed to cancer. Specifically, the invention pertains to a genetically altered non-human animal (most preferably, a mouse), or a cell (either non-human animal or human) in culture, that is defective in at least one of two alleles of a tumor-suppressor gene. The inactivation of at least one of these tumor suppressor alleles results in an animal with a higher susceptibility to tumor induction or other proliferative or differentiative disorders, or disorders marked by aberrant signal transduction, e.g., from a cytokine or

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growth factor. A genetically altered mouse of this type is able to serve as a useful model for hereditary cancers and as a test animal for carcinogen studies. The invention additionally pertains to the use of such non-human animals or animal cells, and their progeny in research and medicine.

Furthermore, it is contemplated that cells of the transgenic animals of the present invention can include other transgenes, e.g., which alter the biological activity of a second tumor suppressor gene or an oncogene. For instance, the second transgene can functionally disrupt the biological activity of a second tumor suppressor gene, such as p53, p73, DCC, p21cip1, p27kip1, Rb, Mad or E2F. Alternatively, the second transgene can cause overexpression or loss of regulation of an oncogene, such as ras, myc, a cdc25 phosphatase, Bcl-2, Bcl-6, a transforming growth factor, neu, int-3, polyoma virus middle T antigen, SV40 large T antigen, a papillomaviral E6 protein, a papillomaviral E7 protein, CDK4, or cyclin D1.

A preferred transgenic non-human animal of the present invention has germline and/or somatic cells in which one or more alleles of a gene are disrupted by a chromosomally incorporated transgene, wherein the transgene includes a marker sequence providing a detectable signal for identifying the presence of the transgene in cells of the transgenic animal, and replaces at least a portion of the gene or is inserted into the gene or disrupts expression of a wild-type protein.

Still another aspect of the present invention relates to methods for generating non-human animals and stem cells having a functionally disrupted endogenous gene. In a preferred embodiment, the method comprises the steps of:

- (i) constructing a transgene construct including (a) a recombination region having at least a portion of the gene, which recombination region directs recombination of the transgene with the gene, and (b) a marker sequence which provides a detectable signal for identifying the presence of the transgene in a cell;
- (ii) transfering the transgene into stem cells of a non-human animal;
- (iii) selecting stem cells having a correctly targeted homologous recombination between the transgene and the gene;
- (iv) transfering cells identified in step (iii) into a non-human blastocyst and implanting the resulting chimeric blastocyst into a non-human female; and

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(v) collecting offspring harboring an endogenous gene allele having the correctly targeted recombination.

Yet another aspect of the invention provides a method for evaluating the carcinogenic potential of an agent by (i) contacting a transgenic animal of the present invention with a test agent, and (ii) comparing the number of transformed cells in a sample from the treated animal with the number of transformed cells in a sample from an untreated transgenic animal or transgenic animal treated with a control agent. The difference in the number of transformed cells in the treated animal, relative to the number of transformed cells in the absence of treatment with a control agent, indicates the carcinogenic potential of the test compound.

Another aspect of the invention provides a method of evaluating an antiproliferative activity of a test compound. In preferred embodiments, the method includes contacting a transgenic animal of the present invention, or a sample of cells from such animal, with a test agent, and determining the number of transformed cells in a specimen from the transgenic animal or in the sample of cells. A statistically significant decrease in the number of transformed cells, relative to the number of transformed cells in the absence of the test agent, indicates the test compound is a potential anti-proliferative agent.

The practice of the present invention will employ, unless otherwise indicated, conventional techniques of cell biology, cell culture, molecular biology, transgenic biology, microbiology, recombinant DNA, and immunology, which are within the skill of the art. Such techniques are explained fully in the literature. See, for example, Molecular Cloning A Laboratory Manual, 2nd Ed., ed. by Sambrook, Fritsch and Maniatis (Cold Spring Harbor Laboratory Press:1989); DNA Cloning, Volumes I and II (D. N. Glover ed., 1985); Oligonucleotide Synthesis (M. J. Gait ed., 25 1984); Mullis et al. U.S. Patent No. 4,683,195; Nucleic Acid Hybridization (B.D. Hames & S. J. Higgins eds. 1984); Transcription And Translation (B. D. Hames & S. J. Higgins eds. 1984); Culture Of Animal Cells (R. I. Freshney, Alan R. Liss, Inc., 1987); Immobilized Cells And Enzymes (IRL Press, 1986); B. Perbal, A Practical Guide To Molecular Cloning (1984); the treatise, Methods In Enzymology (Academic Press, Inc., N.Y.); Gene Transfer Vectors For Mammalian Cells (J. H. Miller and M. P. Calos eds., 1987, Cold Spring Harbor Laboratory); Methods In Enzymology, Vols. 154 and 155 (Wu et al. eds.), Immunochemical Methods In Cell And Molecular

Biology (Mayer and Walker, eds., Academic Press, London, 1987); Handbook Of Experimental Immunology, Volumes I-IV (D. M. Weir and C. C. Blackwell, eds., 1986); Manipulating the Mouse Embryo, (Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y., 1986).

As mentioned above, the sequences described herein are believed to have particular utility in regards to colon cancer. However, they may also be useful with other types of cancers and other disease states.

The present invention will now be illustrated by reference to the following examples which set forth particularly advantageous embodiments. However, it should be noted that these embodiments are illustrative and are not to be construed as restricting the invention in any way.

## XI. Examples

A. Identification of differentially expressed sequences.

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# Description of the Libraries

SEQ ID Nos: 1-544 were derived from libraries designated as DE and PA as described below. The DE library is a normalized, colon cancer specific, subtracted cDNA library. The DE library is specific for sequences expressed in colon cancer [proximal and distal Dukes' B, microsatellite instability negative (MSI-)] but not expressed in normal tissues, including normal colon tissue. The PA library is a normalized, colon specific, subtracted cDNA library. The PA library is specific for sequences expressed in normal colon tissue but not expressed in other normal tissues.

## 25 Construction of a colon cancer specific library

A subtracted colon cancer specific library was made by subtracting pooled proximal, stage B, MSI and distal, Stage B, MSI tumor tissue cDNA against a combination of pooled driver normal cDNA made from colon, peripheral blood leukocytes (PBL), liver, spleen, lung, kidney, heart, small intestine, skeletal muscle, and prostate tissue cDNAs. The following RNA samples were obtained from Origene Technologies, Inc., Rockville, Maryland, and were used to synthesize the pooled driver cDNA: #HT-1015 normal colon total RNA, #HT-1005 liver total RNA, #HT-1004 spleen total RNA, #HT-1009 lung total RNA, #HT-1003 kidney total RNA,

#HT-1006 peripheral blood leukocyte total RNA, #HT-prostate total RNA, #HM-1002 heart muscle poly A+ RNA, #HM-1007 intestine poly A+ RNA, and #HM-1008 skeletal muscle poly A+ RNA. First-strand cDNA was prepared for each using 1 microgram of RNA. A biased pool of first-strand cDNA was prepared containing 50% normal colon first-strand cDNA reaction and 5.56% of each of the remaining tissue first-strand cDNA reactions by volume. Eight individual amplification reactions, each containing 1 microliter of the biased first-strand cDNA reaction pool, were performed for 18 cycles. The double stranded cDNA product from all eight amplification reactions were pooled and purified for subsequent use in subtractive hybridization. The colon cancer specific subtracted library was called DE and individual clones derived from this library were referred to with a number prefixed by DE.

Normalized subtracted DE colon cancer specific and pooled normal human tissue specific cDNA libraries (same as components of driver cDNA above) were generated according published procedures (Daitchenko et al., 1996 PNAS 93:6025-6030, Gurskaya et al., 1996 Analytical Biochemistry 240:90-97) using Clontech Laboratories, Inc., PCR-Select cDNA subtraction kit, PT1117-1. A forty-five fold mass excess of driver cDNA (450 nanograms) was used for each subtraction experiment. Subtractive hybridization of tester with driver cDNAs was performed twice, each time for about 8 -12 hours. Subtracted cancer specific DE cDNA was ligated into the pCR2.1-TOPO plasmid vector (Invitrogen Corporation, Carlsbad CA) and chemically transformed into ultracompetent Epicurian E. coli XL10-Gold cells (Stratagene, La Jolla, CA). A reverse library was also constructed wherein the tester and driver samples were switched; this library was designated as MD.

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#### Construction of a normal colon specific library

This normal colon tissue specific library was made using Clontech Laboratories Inc PCR-Select kit, K1804-1, following instructions from the users manual (PT1117-1).

Four, 100 µl, SMART PCR cDNA amplification reactions for each normal, non-cancerous, patient sample, were performed, starting with 1 µl from their respective first strand cDNA reactions. Each sample was amplified for only 18 cycles using the following PCR conditions; 95 C-10 sec, 68 C 5 min. using a 9600 Perkin

Elmer instrument. The following are Bayer Diagnostic sample identification numbers for the cDNA samples that were amplified: NPB(-) 27347, NPB(-)27859, NPB(-)28147, NPB(-)28162, NDB(-)28800, NDB(-)29243, NDB(-)29244 and NDB(-)42472. These are normal colon tissue samples obtained from the same patients providing the proximal stage B MSI - and distal stage B MSI- cancer samples, which were used to prepare the DE library described above. Equal volumes of the eight normal colon cDNAs were pooled. A subtracted normal colon tissue specific library was made by subtracting the normal colon cDNA pool against a combination of pooled driver normal cDNA made from peripheral blood leukocytes (PBL), liver, spleen, lung, kidney, heart, small intestine, skeletal muscle, and prostate tissue cDNAs. The following are the RNA samples that were used to synthesize the pooled driver cDNA: #HT-1005 liver total RNA, #HT-1004 spleen total RNA, #HT-1009 lung total RNA, #HT-1003 kidney total RNA, #HT-1006 peripheral blood leukocyte total RNA, #HT-prostate total RNA, #HM-1002 heart muscle poly A+ RNA, #HM-1007 intestine poly A+ RNA, and #HM-1008 skeletal muscle poly A+ RNA. First-strand cDNA was prepared for each using 1 microgram of RNA. A pool of first strand cDNA reactions was then made consisting of equal volumes of the nine driver tissue first-strand cDNA reactions. Eight individual amplification reactions, each containing 1 microliter of the first-strand cDNA reaction pool, were performed for 18 cycles. The double stranded cDNA product from all eight amplification reactions was pooled and purified for subsequent use in subtractive hybridization. The normal colon tissue specific subtracted library was called PA and individual clones derived from this library were referred to with a number prefixed by PA.

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The normalized subtracted PA normal colon specific cDNA library and a subtracted normal human tissue specific cDNA library, consisting of the human tissues listed above were generated according published procedures (Daitchenko et al., 1996 PNAS 93:6025-6030, Gurskaya et al., 1996 Analytical Biochemistry 240:90-97) using Clontech Laboratories, Inc., PCR-Select cDNA subtraction kit, PT1117-1. Library construction and cloning were carried out as described above for the colon cancer specific library. Out of the 1152 clones that were analyzed for differential expression, approximately 69% were differentially expressed.

Each EST isolated from each of the above libraries represents a sequence from a partial mRNA transcript, since the cDNA used for making the subtracted library

was restricted with RsaI, a four base cutter restriction endonuclease that generates fragments with an average size of about 600 base pairs.

## Validation of differential expression in colon cancer

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To validate that the differentially expressed sequences found in this library were specific to colon cancer, the clones were screened with cDNAs prepared from a colon cancer specific library, Delaware (DE), and a normal tissue specific library Maryland (MD).

cDNA clones were analyzed for differential expression following the procedure developed by von Stein et al., 1997, Nucleic Acids Research 25(13):2598-2602 and using probes synthesized according to a published method (Jin et al., 1997, Biotechniques 23:1083-1086). Out of the 1248 clones that were analyzed for differential expression approximately 83% were differentially expressed.

## Sequencing and analysis of differentially expressed clones

The nucleotide sequence of the inserts from clones shown to be differentially expressed was determined by single-pass sequencing from either the T7 or M13 promoter sites using fluorescently labeled dideoxynucleotides via the Sanger sequencing method. Sequences were analyzed according to methods described in the text (XI., Examples; B. Results of Public Database Search).

Each nucleic acid represents sequence from at least a partial mRNA transcript. The nucleic acids of the invention were assigned a sequence identification number (see attachments). The nucleic acid sequences are provided in the attached Sequence Listing.

An example of an experiment to identify differentially expressed clones is shown in the Figure, "Differential Expression Analysis". The inserts from subtracted clones were amplified, electrophoresed, and blotted on to membranes as described above. The gel was hybridized with RSA1 cut DE and MD cDNA probes as described above.

In the Figure, individual clones are designated by a number at the top of each lane; the blots are aligned so that the same clone is represented in the same vertical lane in both the upper ("Cancer Probe") and lower ("Normal Probe") blot. Lanes labeled "O" indicate clones that are overexpressed, i.e., show a darker, more prominent band in the upper blot ("Cancer Probe") relative to that observed, in the

same lane, in the lower blot ("Normal Probe"). The Lane labeled "U" indicates a clone that is underexpressed, i.e., shows a darker, more prominent band in the lower blot ("Normal Probe") relative to that observed, in the same lane, in the upper blot ("Cancer Probe"). The lane labeled "M", indicates a clone that is marginally overexpressed in cancer and normal cells.

#### B. Results of Public Databases Searches

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The nucleotide sequence of SEQ ID Nos. 1-544 were aligned with individual sequences that were publicly available. Genbank and divisions of GenBank, such as dbEST, CGAP, and Unigene were the primary databases used to perform the sequence similarity searches. The patent database, GENESEQ, was also utilized.

A total of 544 sequences were analyzed. The sequences were first masked to identify vector-derived sequences, which were subsequently removed. The remaining sequence information was used to create the Sequence Listing (SEQ ID Nos. 1-544). Each of these sequences was used as the query sequence to perform a Blast 2 search against the databases listed above. The Blast 2 search differs from the traditional Blast search in that it allows for the introduction of gaps in order to produce an optimal alignment of two sequences.

A proprietary algorithm was developed to utilize the output from the Blast 2 searches and categorize the sequences based upon high similarity (e value < 1e-40) or identity to entries contained in the GenBank and dbEST databases. Three categories were created as follows: 1) matches to known human genes, 2) matches to human EST sequences, and 3) no significant match to either 1 or 2, and therefore a potentially novel human sequence.

Those skilled in the art will recognize, or be able to ascertain, using not more than routine experimentation, many equivalents to the specific embodiments of the invention described herein. Such specific embodiments and equivalents are intended to be encompassed by the following claims.

All patents, published patent applications, and publications cited herein are incorporated by reference as if set forth fully herein.

# TABLE 1

			050 10 110	-1	Tissue Probe
SEQ ID NO	clone name	Tissue Probe	SEQ ID NO	cione name de0079t7	N
1	de0020t7	U	53 54	de007917 de0085t7	N
2	de0041t7	N U	55	de0089t7	N
3	de0056t7		56	de0095t7	N N
4	de0064t7	N	57	de0093t7	N
5	de0092t7	U	57 58	de009517 de0105t7	N
6	de0142t7	N	59	de010317 de0112t7	N
7	de0153t7	M	60	de011217 de0114t7	N
8	de0163t7	U	61	de011417 de012117	N
9	de0188t7	N	62	de012117 de012217	N
10	de0190t7	U	63	de012217 de0124t7	N
11	de0201t7	M			M
12	de0225t7	U	64	de0139t7 de0143t7	N N
13	de0246t7	Ü	65 66	de014317 de0166t7	Ü
14	de0257t7	N	66 67		N
15	de0285t7	0	67	de0168t7	
16	de0529t7	U	68	de0171t7	N
17	de0629t7	U	69	de0178t7	N
18	de072717	0	70	de0180t7	0
19	de0787t7	U	71	de0181t7	N
20	de0810t7	N	72	de0199t7	N
21	de0833t7	N	73	de0200t7	N
22	pa0107t7	U	74	de0202t7	N
23	pa0130t7	U	75	de0205t7	N
24	pa0149t7	U	76	de0207t7	U
25	pa0185t7	Ų	77	de0212t7	N
26	pa0203t7	U	78	de0217t7	N
27	pa0277t7	U	79	de0220t7	U
28	pa0287t7	U	80	de0228t7	N
29	pa0293t7*	U	81	de0236t7	0
30	pa0341t7	U	82	de0243t7	N
31	pa0357t7	N	83	de0253t7	0
32	pa0361t7	U	84	de0258t7	N
33	pa0404t7	U	85	de0259t7	N
34	pa0408t7	U	86	de0262t7	N
35	pa0425t7	N	87	de0270t7	N
36	de0001t7	N	88	de0275t7	N
37	de0002t7	N	89	de0287t7	N
38	de0036t7	N	90	de0288t7	N
39	de0038t7	M	91	de0306t7	N
40	de0040t7	N	92	de0490t7	N
41	de0043t7	0	93	de0501t7	M
42	de0044t7	N	94	de0516t7	N
43	de0045t7	N	95	de0589t7	N
44	de0050t7	N	96	de0596t7	U
45	de0052t7	N	97	de0600t7	N
46	de0054t7	N	98	de0609t7	U
47	de0055t7	N	99	de0611t7	N
48	de0059t7	0	100	de0617t7	U
49	de0060t7	N	101	de0633t7	N
50	de0063t7	U	102	de0643t7	N
51	de0066t7	Ō	103	de0647t7	М
52	de006717	0	104	de0652t7	N

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105	de0666t7	N	161	pa0405t7	N
105	de0695t7	Ü	162	pa0406t7	N
107	de0705t7	N	163	pa0409t7	Ü
107	de0706t7	M	164	pa0411t7	N
109	de0708t7	N	165	pa0417t7	N
110	de0724t7	N N	166	pa0421t7	Ü
111	de0735t7	N	167	pa0429t7	Ü
112	de0740t7	N	168	pa0432t7	Ū
113	de0742t7	N	169	de0004t7	Ū
114	de0747t7	N	170	de0008t7	ND
115	de0764t7	N N	171	de0009t7	ND
116	de0777t7	Ö	172	de0010t7	ND
117	de077117	N	173	de0011t7	ND
118	de0793t7	ΰ	174	de0012t7	ND
119	de079317 de079417	N	175	de0013t7	ND
120	de079417 de079817	N	176	de0014t7	ND
121	de0800t7	Ö	177	de0016t7	ND
122	de0800t7 de0816t7	N	178	de0017t7	ND
	de0818t7	N	179	de0018t7	M
123		N	180	de0019t7	ND
124	de0835t7	Ü	181	de0023t7	0
125	pa0078t7	N	182	de0023t7	N
126	pa0080t7	U	183	de0029t7	ND
127	pa0088t7	U	184	de002317	ND
128	pa0089t7	Ü	185	de0032t7	ND
129	pa0095t7	U	186	de0032t7	O
130	pa0158t7	Ü	187	de0033t7	ND
131	pa0159t7 pa0187t7	N	188	de0035t7	ND
132 133	pa010717 pa0190t7	U	189	de0042t7	ND
134	pa019017 pa019217	Ü	190	de0047t7	ND
135	pa019207 pa0209t7	Ü	191	de0048t7	N
136	pa0205t7 pa0215t7	N	192	de0049t7	ND
137	pa0218t7	N N	193	de0051t7	0
138	pa021017	N	194	de0053t7	ND
139	pa0238t7	N	195	de0065t7	ND
140	pa0230t7	Ü	196	de0068t7	N
141	pa024517	N	197	de0069t7	ND
142	pa025017 pa025817	ΰ	198	de007117	N
143	pa0233t7	N	199	de0071t7	ND
144	pa0283t7	N	200	de0076t7	Ü
145	pa0295t7	N	201	de0077t7	ND
146	pa029317 pa0309t7	Ü	202	de0078t7	ND
147	pa030317	N	203	de0080t7	ND
148	pa031417 pa031717	N	204	de0082t7	ND
149	pa031717	N	205	de0086t7	ND
150	pa031317	N	206	de0087t7	ND
151	pa032317 pa033317	N	207	de0088t7	ND
152	pa0336t7	N N	208	de0093t7	N
153	pa0353t7	N	209	de0094t7	ND
154	pa0363t7	N	210	de0097t7	0
155	pa0364t7	N	211	de0098t7	ND
156	pa0366t7	Ü	212	de0100t7	ND
157	pa038217	N	213	de010117	ND
158	pa0383t7	N	214	de0102t7	ND
159	pa0388t7	N	215	de0106t7	ND
160	pa0389t7	N	216	de0109t7	Ü
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			070		
217	de0110t7	N	273	de0214t7	ND
218	de0111t7	N	274	de0215t7	ND
219	de0113t7	ND	275	de0218t7	ND
220	de0115t7	0	276	de0221t7	ND
221	de0117t7	ND	277	de0223t7	0
222	de0118t7	U	278	de0227t7	ND
223	de0119t7	ND	279	de0229t7	0
224	de0123t7	ND	280	de0230t7	ND
225	de0125t7	ND	281	de0232t7	ND
226	de0126t7	ND	282	de0234t7	ND
227	de0129t7	ND	283	de0235t7	ND
228	de0130t7	U	284	de0237t7	ND
229	de0131t7	0	285	de0238t7	ND
230	de0132t7	ND	286	de0239t7	N
231	de0134t7	0	287	de0241t7	N
232	de0135t7	ND	288	de0242t7	0
233	de0137t7	M	289	de0244t7	N
234	de0138t7	ND	290	de0247t7	Ö
235	de0140t7	ND	291	de0252t7	ND
236	de014117	ND	292	de0255t7	N
237	de0145t7	ND	293	de0256t7	ND
238	de0146t7	0	294	de0260t7	N
239	de0148t7	ND	295	de0261t7	N
240	de0149t7	ND	296	de0263t7	N
241	de015117	0	297	de0264t7	, ND
242	de0152t7	ND	298	de0265t7	ND
243	de0154t7	ND	299	de0266t7	0
244	de0156t7	ND	300	de0267t7	N
245	de0157t7	U	301	de0268t7	ND
246	de0158t7	ND	302	de0272t7	ND
247	de0159t7	N	303	de0273t7	ND
248	de0162t7	ND	304		
				de0274t7	N
249	de0169t7	U	305	de0276t7	0
250	de0170t7	0	306	de0277t7	M
251	de0174t7	ND	307	de0279t7	N
252	de0176t7	ND	308	de0280t7	ND
253	de0177t7	0	309	de0281t7	N
254	de0182t7	ND	310	de0282t7	ND
255	de0183t7	ND	311	de0284t7	ND
256	de0184t7	ND	312	de0286t7	ND
257	de0186t7	ND	313	de0339t7	ND
258	de0187t7	M	314	de0483t7	ND
259	de0189t7	ND	315	de0484t7	М
260	de0191t7	M	316	de0491t7	ND
261	de0192t7	ND	317	de0499t7	ND
262	de0193t7	ND	318	de0507t7	М
263	de0195t7	N	319	de0511t7	0
264	de0196t7	N	320	de0519t7	ND
265	de0197t7	N	321	de0520t7	N
266	de0198t7	ND	322	de0522t7	ND
267	de0203t7	ND	323	de052417	M
268	de0208t7	ND	324	de0530t7	ND
269	de0209t7	N	325	de0531t7	ND
270	de0210t7	N	326	de0532t7	M
271	de0211t7	ND	327	de0534t7	N N
272	de021117	ND	328	de0542t7	ND
212	0002100	NU	320	UCUJ4217	עטו

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329	de0556t7	M		385	de0707t7	0
330	de0557t7	ND		386	de0709t7	0
331	de0559t7	U		387	de0710t7	ND
332	de0562t7	ND		388	de0712t7	N
333	de0566t7	U		389	de0715t7	ND
334	de0567t7	N		390	de0719t7	N
335	de0568t7	ND		391	de0722t7	ND
336	de0570t7	ND		392	de0723t7	ND
337	de057117	ND		393	de0725t7	N
338	de0574t7	ND		394	de0728t7	ND
339	de0581t7	ND		395	de0729t7	ND
340	de0583t7	U		396	de0731t7	ИD
341	de0587t7	ND		397	de0732t7	ND
342	de0588t7	ND		398	de0737t7	ND
343	de0591t7	ND		399	de0739t7	М
344	de0592t7	ND		400	de0741t7	ND
345	de0597t7	U		401	de0744t7	N
346	de0598t7	ND		402	de0746t7	ND
347	de0599t7	ND		403	de0749t7	N
348	de0602t7	N		404	de0750t7	ND
349	de0605t7	ND		405	de0756t7	ND
350	de0608t7	ND		406	de0759t7	ND
351	de0610t7	ND		407	de0761t7	0
352	de0616t7	0		408	de0762t7	ND
353	de0619t7	U		409	de0766t7	ND
354	de0620t7	ND		410	de0768t7	U
355	de0622t7	ND		411	de0769t7	ND
356	de0623t7	ND		412	de0772t7	ND
357	de0624t7	О		413	de0776t7	ND
358	de0625t7	. ND	68	414	de0779t7 ~	ND
359	de0628t7	ND		415	de0785t7	ND
360	de0630t7	ND		416	de0786t7	ND
361	de0631t7	ND		417	de0788t7	ND
362	de0632t7	N		418	de0789t7	ND
363	de0634t7	ND	**	419	de0792t7	ND
364	de0639t7	ND .		420	de0796t7	ND
365	de0642t7	ND		421	de0797t7	ND
366	de0649t7	, ND	4.4	422	de0801t7	0
367	de0650t7	N		423	de0804t7	ND
368	de0656t7	N		424	de0805t7	ND
369	de0657t7	ND		425	de0806t7	ND
370	de0660t7	ND		426	de0807t7	N
371	de0661t7	0		427	de0811t7	0
372	de0662t7	0		428	de0812t7	ND
373	de0664t7	ND		429	de0817t7	N
374	de0665t7	ND		430	de0820t7	ND
375	de0667t7	ND		431	de0821t7	ND
376	de0669t7	ND		432	de0822t7	ND
377	de0676t7	ND		433	de0823t7	N
378	de0686t7	N		434	de0824t7	N
379	de0687t7	ND		435	de0825t7	ND
380	de0689t7	N		436	de0826t7	ND
381	de0691t7	М		437	de0827t7	ND
382	de0693t7	ND		438	de0829t7	ND
383	de0703t7	ND		439	de0830t7	ND
384	de0704t7	М		440	de0837t7	N

441	de0840t7	ND	497	pa0240t7	ND
442	de0848t7	ND	498	pa0252t7	
443	pa0079t7	N	499	pa0260t7	
444	pa0081t7	ND	500	pa0261t7	_
445	pa0082t7	ND	501	pa0262t7	ND
446	pa0083t7	ND	502	pa0264t7	N
447	pa0084t7	ND	503	pa0265t7	N
448	pa0085t7	ND	504	pa0268t7	ND
449	pa0086t7	M	505	pa0276t7	ND
450	pa0090t7	N	506	pa0279t7	ND
451	pa0091t7	ND	507	pa0280t7	ND
452	pa0092t7	N	508	pa0282t7	ND
453	pa0096t7	ND	509	pa0285t7	ND
454	pa0100t7	ND	510	pa0299t7	ND
455	pa0101t7	U	511	pa0300t7	Ü
456	pa0103t7	ND	512	pa0301t7	ND
457	pa0104t7	ND	513	pa0302t7	ND
458	pa0114t7	ND	514	pa0305t7	N
459	pa0115t7	ND	515	pa0306t7	ND
460	pa0118t7	ND	516	pa0307t7	ND
461	pa0120t7	ND	517	pa0311t7	ND
462	pa0129t7	ND	518	pa0316t7	ND
463	pa0131t7	U	519	pa0318t7	ND
464	pa0133t7	ND	520	pa0321t7	М
465	pa0135t7	N	521	pa0325t7	N.
466	pa0140t7	0	522	pa0326t7	ND
467	pa0142t7	ND	523	pa0332t7	ND
468	pa0143t7	ND	524	pa0339t7	ND
469	pa0146t7	ND	525	pa0346t7	0
470	pa0147t7	ND	526	pa0349t7	ND
471	pa0148t7	ND	527	pa0351t7	Ü
472	pa0151t7	ND	528	pa0355t7	ND
473	pa0157t7	ND	529	pa0358t7	ND
474	pa0164t7	ND	530	pa0360t7	N
475	pa0167t7	N	531	pa0362t7	ND
476	pa0171t7	U	532	pa0368t7	U
477	pa0174t7	ND	533	pa0369t7	ND
478	pa0175t7	ND	534	pa0373t7	ND
479	pa0179t7	N	535	pa0380t7	ND
480	pa0182t7	ND	536	pa0393t7	ND
481	pa0184t7	ND	537	pa0395t7	ND
482	pa0186t7	U	538	pa0396t7	ND
483	pa0189t7	ND	539	pa0397t7	ND
484	pa0207t7	ND	540	pa0410t7	N
485	pa0210t7	ND	541	pa0415t7	ND
486	pa0212t7	ND	542	pa0416t7	ND
487	pa0214t7	ND	543	pa0424t7	ND
488	pa0216t7	ND	544	pa0430t7	ND
489	pa0217t7	М	•	pac 10011	IID
490	pa0219t7	N	* In the provi	sional annlic	ation (60/098,639
491	pa0223t7	ND	August 31 1	998, clone F	A0293t7 was labe
492	pa0224t7	ND	clone PA002	3t7 in error	That mistake has
493	pa0228t7	ND			the accurate clone
404	02022017	11			2000, 010 010116

494

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pa0229t7

pa0231t7

pa0232t7

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ND

ND

# Table 2 "Novel" Region 2

		"Novel" Region 1	"Novel" Region 2					
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9		S		9	က	က္	4	2	4		ဖွ		4	'n		ίÖ	ίū	ίδ	Ņ	2	~	7	'n	æ	6		4	۰	0
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105.00	106.00	107.00	108.00	110.00	113.00	116.00	117.00	118.00	119.00	120.00	121.00	122.00	123.00	124.00	129.00	132.00	134.00	139.00	140.00	142.00	143.00	148.00	153.00	156.00	157.00	159.00	160.00	161.00	

# TABLE 3

The following list of clones indicates those found in either the DE or PA libraries and the SW480 library

SEQ ID NO clone name  185 de0032t7  186 de0033t7  193 de0051t7  196 de0068t7  240 de0149t7  241 de0151t7  247 de0159t7  72 de0199t7  279 de0229t7  281 de0232t7  283 de0235t7  306 de0277t7  310 de0282t7  318 de0507t7  328 de0559t7  342 de0588t7
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196 de0068t7 240 de0149t7 241 de0159t7 72 de0199t7 279 de0229t7 281 de0232t7 283 de0235t7 306 de0277t7 310 de0282t7 318 de0507t7 328 de0542t7 331 de0559t7
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241 de0151t7 247 de0159t7 72 de0199t7 279 de0229t7 281 de0232t7 283 de0235t7 306 de0277t7 310 de0282t7 318 de0507t7 328 de0542t7 331 de0559t7
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410 de0768t7
427 de0811t7
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470 pa0147t7
481 pa0184t7
493 pa0228t7
494 pa0229t7
140 pa0249t7
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510 pa0299t7
515 pa0306t7
517 pa0311t7
518 pa0316t7
536 pa0393t7
539 pa0397t7
544 pa0430t7

#### We claim:

 An isolated nucleic acid comprising a nucleotide sequence which hybridizes under stringent conditions to a sequence of SEQ ID Nos. 1-35 or a sequence complementary thereto.

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- An isolated nucleic acid comprising a nucleotide sequence at least 80% identical to a sequence corresponding to at least about 15 consecutive nucleotides of one of SEQ ID Nos. 1-35 or a sequence complementary thereto.
- 3. An isolated nucleic acid comprising a nucleotide sequence of SEQ ID Nos. 1-35 or a sequence complementary thereto.
- A nucleic acid according to claim 1, further comprising a transcriptional regulatory sequence operably linked to said nucleotide sequence so as to
   render said nucleotide sequence suitable for use as an expression vector.
  - 5. An expression vector, capable of replicating in at least one of a prokaryotic cell and eukaryotic cell, comprising the nucleic acid of claim 4.
- 20 6. A host cell transfected with the expression vector of claim 5.
  - 7. A transgenic animal having a transgene of the nucleic acid of claim 1 incorporated in cells thereof, which transgene modifies the level of expression of the nucleic acid, the stability of an mRNA transcript of the nucleic acid, or the activity of the encoded product of the nucleic acid.
  - 8. A substantially pure nucleic acid which hybridizes under stringent conditions to a nucleic acid probe corresponding to at least 12 consecutive nucleotides of one of SEQ ID Nos. 1-168 or a sequence complementary thereto.

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 A polypeptide including an amino acid sequence encoded by a nucleic acid of claim 1 or a fragment comprising at least 25 amino acids thereof.

10. A probe/primer comprising a substantially purified oligonucleotide, said oligonucleotide containing a region of nucleotide sequence which hybridizes under stringent conditions to at least 12 consecutive nucleotides of sense or antisense sequence selected from SEQ ID Nos. 1-168.

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- 11. An array including at least 10 different probes of claim 10 attached to a solid support.
- 12. The probe/primer of claim 10, further comprising a label group attached thereto and able to be detected.
  - 13. The probe/primer of claim 12, wherein said label group being selected from radioisotopes, fluorescent compounds, enzymes, and enzyme co-factors.
- 15 14. An antibody immunoreactive with a polypeptide of claim 9.
  - 15. An antisense oligonucleotide analog which hybridizes under stringent conditions to at least 12 consecutive nucleotides of one of SEQ ID Nos. 1-35 or a sequence complementary thereto, and which is resistant to cleavage by a nuclease.
  - 16. A test kit for determining the phenotype of transformed cells, comprising the probe/primer of claim 12, for measuring a level of a nucleic acid which hybridizes under stringent conditions to a nucleic acid of SEQ ID Nos. 1-544 in a sample of cells isolated from a patient.
  - 17. A test kit for determining the phenotype of transformed cells, comprising an antibody specific for a protein encoded by a nucleic acid which hybridizes under stringent conditions to any one of SEQ Nos. 1-544.

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18. A method of determining the phenotype of a cell, comprising detecting the differential expression, relative to a normal cell, of at least one nucleic acid

which hybridizes under stringent conditions to one of SEQ ID Nos. 1-544, wherein the nucleic acid is differentially expressed by at least a factor of two.

- 19. A method for determining the phenotype of cells in a sample of cells from a patient, comprising:
  - i. providing a nucleic acid probe comprising a nucleotide sequence having at least 12 consecutive nucleotides of any of SEQ ID Nos. 1-544;
  - ii. obtaining a sample of cells from a patient;

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- iii. providing a second sample of cells substantially all of which are non-cancerous;
- iv. contacting the nucleic acid probe under stringent conditions with mRNA of each of said first and second cell samples; and
- v. comparing (a) the amount of hybridization of the probe with mRNA of the first cell sample, with (b) the amount of hybridization of the probe with mRNA of the second cell sample, wherein a difference of at least a factor of two in the amount of hybridization with the mRNA of the first cell sample as compared to the amount of hybridization with the mRNA of the second cell sample is indicative of the phenotype of cells in the first cell sample.
- 20. A method of determining the phenotype of a cell, comprising detecting the differential expression, relative to a normal cell, of at least one protein encoded by a nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-544, wherein the protein is differentially expressed by at least a factor of two.
- 21. The method of claim 20, wherein the level of said protein is detected in an immunoassay.
- 22. A method for determining the presence or absence of a nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-168 in a cell, comprising contacting the cell with a probe of claim 10.

23. A method for determining the presence of absence of a polypeptide encoded by a nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-35 in a cell, comprising contacting the cell with an antibody of claim 14.

- 24. A method for detecting a mutation in a test nucleic acid which hybridizes under stringent conditions to a nucleic acid of SEQ ID Nos. 1-544 or a sequence complementary thereto, comprising
  - i. collecting a sample of cells from a patient,
  - ii. isolating nucleic acid from the cells of the sample,
  - iii. contacting the nucleic acid sample with one or more primers which specifically hybridize to a nucleic acid sequence of SEQ ID Nos. 1-544 under conditions such that hybridization and amplification of the nucleic acid occurs, and
  - iv. comparing the presence, absence, or size of an amplification product to the amplification product of a normal cell.
- 25. A method for identifying an agent which alters the level of expression in a cell
  20 of a nucleic acid which hybridizes under stringent conditions to one of SEQ ID
  Nos. 1-544 or a sequence complementary thereto, comprising
  - i. providing a cell;
  - ii. treating the cell with a test agent;
  - iii. determining the level of expression in the cell of a nucleic acid which hybridizes under stringent conditions to one of SEQ ID Nos. 1-544 or a sequence complementary thereto; and
  - iv. comparing the level of expression of the nucleic acid in the treated cell with the level of expression of the nucleic acid in an untreated cell, wherein a change in the level of expression of the nucleic acid in the treated cell relative to the level of expression of the nucleic acid in the untreated cell is indicative of an agent which alters the level of expression of the nucleic acid in a cell.

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26. A pharmaceutical composition comprising an agent identified by the method of claim 25.

- A pharmaceutical composition comprising a nucleic acid which includes a
   nucleotide sequence which hybridizes under stringent conditions to one of
   SEQ ID Nos. 1-544 or a sequence complementary thereto.
- 28. A pharmaceutical composition comprising a polypeptide encoded by a nucleic acid which includes a nucleotide sequence that hybridizes under stringent conditions to one of SEQ ID Nos. 1-544 or a sequence complementary thereto.
  - An isolated nucleic acid comprising a portion of a nucleotide sequence of SEQ
     ID Nos. 36-168 or a sequence complementary thereto.
- 15 30. A gene which hybridizes to one of SEQ ID Nos. 1-35.

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- 31. A method for detecting cancer in which one or more of SEQ ID Nos. 1-544 are used as probes, said method comprising:
  - i. collecting a sample of cells from a patient,
  - ii. isolating nucleic acid from the cells of the sample,
  - iii. contacting the nucleic acid sample with one or more primers which specifically hybridize to a nucleic acid sequence of SEQ ID Nos. 1-544 under conditions such that hybridization and amplification of the nucleic acid occurs, and
- iv. comparing the presence, absence, or size of an amplification product to the amplification product of a normal cell.
  - 32. A method of claim 31 in which said cancer is colon cancer.
  - 33. A method for detecting cancer in a patient sample in which an antibody to a protein encoded by SEQ ID Nos. 1-544 is used to react with proteins in said sample.

34. A method of claim 33 in which said cancer is colon cancer.

# Differential Expression Analysis

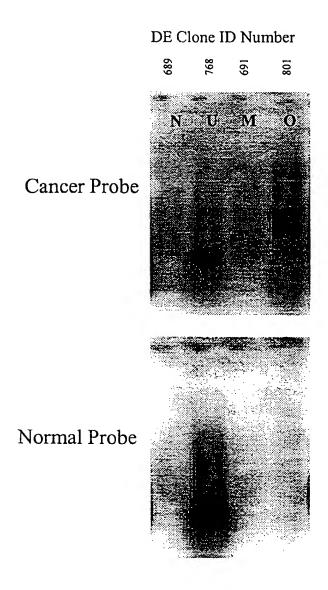


Fig. 1

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                                                                             120
agtaataaca gcatgacttc cctaagatct gattcagaga attgaaatat gccctgagaa
                                                                             180
aacataagag gtttttctgg agaagtgtcc caagggtaat attaattgtt caaggatgtt
                                                                             240
teggaaaaag ttgcaatcat caetgtggca aatgaateta gggagaggaa geatgagtta
                                                                             300
tttaatgtca gttactcctt tccgtaggtt tttgcctttt tttggacttt acacacagcc
                                                                             360
catttgctat gaaactatca gctcaaatag cangctttca ngcaggccaa caatggcaga
                                                                             420
ctgcattctt nctactttnt ccaatcatat ttatcaagtc ccattgggag aatactttca
                                                                             480
gtagngctca aantacccgc ntncaattgg aactgcangg aaccnttcag aaataacnct
                                                                             540
tnaagaaaga aataaccett canggaanac cetttnggnt teactetann tggggttnac
                                                                             600
aagaaa
                                                                             606
      <210> 7
      <211> 620
      <212> DNA
      <213> Homo sapiens
```

```
<220>
      <221> misc_feature
      <222> (1) ... (620)
      <223> n = A, T, C \text{ or } G
      <400> 7
eegtggeeee ggeegaaggt eenetggane eeegggtggt aattggetgg aggtaaatgg
                                                                         60
tggtaangta ttaaccattt ctatggaaat gnccctttgg ggccctcctg gattttaaaa
                                                                        120
tggtcccctg gtttggacnt ttctattaaa gaaatggnca ttttacctaa aatgccnggt
                                                                        180
ctaccttatt aaagancaaa tngnntattn gaccttaaaa taggcatttt tcctaatcat
                                                                        240
aatctggccg gcttaacccc aatcaagata attgggtgcc cnttatgaat ttgaagttag
                                                                        300
tgatageete ettgtaaggt getaceetna tggggataga gaceccaget actantaatt
                                                                        360
ngggaaaatg gttaaggtat ttgggaaaag tactctttta aaaacatatt ggccacagaa
                                                                        420
ancctagget gaattaenng gattgataat tttgnaanta atttentana atgggennge
                                                                        480
tggatgaaaa aatggcctcc tcnttttccc tggaaccagc ngctttttgc ctaaacntta
                                                                        540
ncctttttaa gttgaaccta gggaccacct aatnggente acaatteeet tttteettte
                                                                        600
ctttttttt gcccaagggn
                                                                        620
      <210> 8
      <211> 263
      <212> DNA
      <213> Homo sapiens
      <400> 8
gcgtgggtcg cggccgaggt accaettttc ttattgcaac tcaacaagtg gcaattggtg
atgaaaagtc aagtggggaa cccagtctgt ggggaacaaa tggaataact tacctgtcac
                                                                        120
cttgtctaac cgggatgcaa atcctcaagt ggtattaaaa agcatacagt gttttataac
                                                                        180
tgtagttgtg tggaaagtaa ctggtctcca agaacagaaa ttactcagcg cacttgggtg
                                                                        240
aatgcccaag aaataatact tgt
                                                                        263
      <210> 9
      <211> 590
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(590)
      <223> n = A,T,C or G
      <400> 9
acaacagggt tettgcatca agettcatge ttteccagae atttactcaa gggaacgtgg
                                                                        60
gagagggagg aggagggg gagctgggag tgataagcag atgttacaca tgtttttcct
ggaaagatca ccccactttt tctaatttcc cagaattaaa agaatgtatt ttatctgtat
                                                                        120
                                                                        180
taccatggaa attactagta acactggatt tttttccctc ttttctaaag tttccaaaaa
                                                                       240
ctttcaaaag tgttcaaaga aattttcttg aacaatttta atatgtttga tttctcattt
                                                                        300
ggggctggaa tatttgtatt ctttttaatt tttttacttc atttattaga agaagtttct
                                                                       360
420
gatatatgta gatatatnga tttatgtcnc aatatcactn taaggcattc ttcttccatc
                                                                       480
cttttatatc tncccaaact ggtntnatgg gacctgtcct gcctgtaggt aaaanccttn
                                                                       540
taattteeet gaaaggetae enetttetan ggggneaace aattgggagn
                                                                       590
```

<210> 10

```
<211> 609
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (609)
      <223> n = A, T, C or G
      <400> 10
cgaggtacgt ttttcaatgt tttaaaaaat tgaaagggag tataatgttt cataacacat
                                                                      60
gggaaattat gtgcagctca aatttcaagt atccataaat aaagttttat tggaacacag
                                                                     120
ctacgctcac tcattagata ttgtctatgg ctgtttttgt gcaaaatggc aganttgggt
                                                                     180
tragagttag caaragagag cttgtagcct graagcctag agtatttact atrtggattt
                                                                     240
ctacagaaaa aaaaaattat tgccccctgc catacagtct gactgatagc ctgagaaagt
                                                                     300
atgcattaaa agaaagttac ctaccetgac cecatgagaa tgaatttgaa aagaacenag
                                                                     360
atgtggtaga agcagatagg ctatgaaagt ttcagaaggg tancatcact qtqqqcnaqq
                                                                     420
atattcaaga aaagacttca nggaaaatgt nggggtttga actggncttg agtaggagtt
                                                                     480
naacttangg gaactggntt taggtngcca ctttaaggct gtcaaanatc atggcccaac
                                                                     540
atteantitg geceaaatte eccangngee ttaaaaattt ggacatgget tgggttgggg
                                                                     600
gncaccctt
                                                                     609
      <210> 11
      <211> 578
      <212> DNA
      <213> Homo sapiens
     <220>
      <221> misc_feature
      <222> (1)...(578)
     <223> n = A, T, C \text{ or } G
     <400> 11
acgcgggatg tagtcagagg aggccctgac atctgcaggg cagcatgggt caaaccaaaa
                                                                      60
agacttttct gaggttgggc geagtggete acgeetgtaa teccaacact ttggaaggee
                                                                     120
agtaggggcg gatcacctga ggtcaggaga ttcgagacca tcctggctaa cacggtgaaa
                                                                     180
ccccatctct actaaaaaa atacgaaaaa aattagccag gcgtggtgac gggtgcctgt
                                                                     240
agtcccagct actagggagg ctgaggcagg agaatggtgt gaacccggga ggcagagctt
                                                                     300
gcagtgagcc gagatcaggc cactgcactc cagcctgggc cacaagagcg agactctgtc
                                                                     360
420
ggaatgctgg tgcatgcctg tatcccaact ctcaggaggt tgaagcagga gaatcacctt
                                                                     480
gacccatnag caatgitcat gaacttagne engeentgga etteaneaag geacegagta
                                                                     540
aganttentt tnaaaaaaaa aannnaaaaa aaagteet
                                                                     578
     <210> 12
     <211> 581
     <212> DNA
     <213> Homo sapiens
     <220>
     <221> misc feature
     <222> (1)...(581)
```

<223> n = A,T,C or G

```
<400> 12
actittttt tttttttt tttttttt gggacggagt cttgctctgt tgcccaggct
                                                                              60
ggagtgcagt ggcgccatct tggctcacta caagctccgc ctcccgggtt cacaccattc
                                                                             120
tectgtetta geeteecage geeegeeace geaccegget aattititgt attittagta
                                                                             180
gagacagggt ttcaccatgt tagccaggat ggtctcgatc tcctgacctc gtggcccacc tgccttggcc tccaaaagtg ctggaattac agtcgtgagc caccacgcc ggcctaaacc atttctcttg acaacactct ggattttatt tctggccaga taccatttat caattttacc
                                                                             240
                                                                             300
                                                                             360
atcaagaata agataatcaa aataataatc aagttttata ttagacttat gaagattctt
                                                                             420
geacetttga aattacaget ateteactag trnattetee teteteatat tttattaeng
                                                                             480
acntecagga agacaaccaa cacetttaaa agttggetga geattttta nggagaccet
                                                                             540
taggtaanag ggncetngge gggaaceeet taggggnaat n
                                                                             581
       <210> 13
       <211> 607
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
       <222> (1) ... (607)
      <223> n = A, T, C \text{ or } G
       <400> 13
ggtactggaa caactataag acccctgttc agattaagga atttggtgca gtttcaaaag
                                                                              60
tagacttttc tectcagect ccatataatt atgetgteac agetteetea agaatteaca
                                                                             120
tttatggccg atactcccaa gaacctataa aaaccttttc tcgatttaaa gacacagcat
                                                                             180
actgtgctac ttttcgacaa gatggtagat tgcttgtggc tggcagtgaa gatggtggag
                                                                             240
ttcaactttt tgatataagt gggagggctc ccctcaggca gtttgaaggc catacaaaag
                                                                             300
cagttcatac agtagatttt acagctgaca aatatcacgt ggtctctggg gctgatgatt
                                                                             360
atacagttaa attatgggat attccaaact ccaaagaaat ttttgacatt taaaggaaca
                                                                             420
ctctgattat gtgangtgtg gatgtgctag caaactttaa tccggatctc tttataacca
                                                                             480
gggacatatg atcatactgn gaagatgttg gatgenegaa cenattgaaa agtggtettt
                                                                             540
ccgttgagca tggccnncag tngaaantgn cctacttttc cccttggaag gctttggggt
                                                                             600
annangg
                                                                             607
      <210> 14
      <211> 599
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (599)
      <223> n = A, T, C \text{ or } G
      <400> 14
ggtactttca aaatataaca attttcgttc tcccatataa cagggggcta acaagaaaac
                                                                              60
caaaaataaa taaaaagaga aaatttaaaa ataagtaaaa aataaaaaaa tatttttaaa
                                                                             120
aagcagcctg ggcaagagaa gtgggtgggt ttaggagaat ccctttcgaa aaattcagag
                                                                             180
cattattatt aatcgttctt aaattaaatg cagggccaag catgctgcac gtggaatctg
                                                                             240
gacaattttt tgataaactt taaggctgct aaataattta cagaaactgt gaatgcattt
                                                                             300
tcattttacg aggcaaaaga gaaaatattc aagattgcat agcaatttta ttttttgaaa
                                                                             360
```

```
tggttatcct aaagaatttc cttaaattca gattttgcaa aattcctact ctncaagtca
                                                                        420
tcaagtgaac actaaaagca actttctcgt gaatcagtgg acttttacga ggcatqcatt
                                                                        480
tttcataaat ctaggccaag tgacctaatt gngattaaat cttaatcatc ctgngattct
                                                                        540
ggctattaan atgggtttaa ancngtaaaa atnctttnaa aaagccgtta ctinccgan
                                                                        599
      <210> 15
      <211> 457
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(457)
      \langle 223 \rangle n = A,T,C or G
      <400> 15
ggtacttttt ttttttttt ttttttttc gaaatgaaca aatatttatt tatcttttat
                                                                         60
aacaagtaag gcaatgttgc ttaaaggaag acaaacaaac ataaaagatt ccqttqacaa
                                                                        120
tgcatttttt catnigttcg gcacaatgct titgtcataa tggagatgtg acagcaaact
                                                                        180
ttccaggaca ttcagtcttc ggnggcagca cttagggcan atgactggcc gctcaaattc
                                                                        240
tctatnttgt ttcaggacag tggaaaagct tatanatgag gccaaagcac caggtaggtg
                                                                        300
gaaggttett gtateggtte gaaceeegae agegegeeaa cagacaacae naggeagtgg
                                                                        360
ggagcaacat gctgttttaa tgancgcctg ggtgcangcg tgctgaggct gaaaatggca
                                                                        420
taaccccgc gtcctgccng gcgggcgttc aaanggn
                                                                        457
      <210> 16
      <211> 643
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (643)
      \langle 223 \rangle n = A,T,C or G
      <400> 16
ggtacaatct agctgaaatc atatacaagt aagtaggtgt ggacttttac tgctgagcta
aagtttatgt ttatatatgt tttattcttt aagctaaaca aacattcaga taacattcta
                                                                        120
tgcatttttt gaagcatagg gttagtaatg aggacttaga ttttttaatt aaacaattca
                                                                        180
gtaactatat aaaaagaaaa ggagtccctt atgaataaat attaaaatta aaagaaatag
                                                                        240
gcaactataa aagtaagtat ttttaataat ggcattgatt ttaqtaaqaa atcaattaqq
                                                                        300
ctgggctgga aagaaaaact ggcttaatat aaagtagttt taatatggca aatattette
                                                                        360
ttaaaattgn ggccctggaa tatcatttct gcctattgct gatgctaagg natcaactgn
                                                                        420
gccaagtatt gggctgntcc acaggtggga angagtagca acattttgng gattttttt
                                                                        480
ttttttaaa accggagaat acccggccag gggntcaagn ctgnatccac antttgggag
                                                                        540
nttagccgga naancettgg anceggagna aaggttnaan gagncaaaat gngccatggn
                                                                        600
ttccanctgg ggacccgggg gnaactcttt taaaccnaaa aat
                                                                        643
      <210> 17
      <211> 336
      <212> DNA
      <213> Homo sapiens
```

```
<220>
      <221> misc feature
      <222> (1) ... (336)
      <223> n = A, T, C \text{ or } G
      <400> 17
ggtactttga taaatgtaga aagattattt aattctggct tggtaccgtg gctcatgcct
                                                                         60
ataatcccag cacttcagga ggctgaggtg ggtggatcac ttgagctcag gagtttgaga
                                                                        120
ccaggcgaaa ccctgtctcc acaaaaaatg caaaaattgc tggacatggt ggcacatgcc
                                                                        180
tgtagtccca gctacttgga aggctgaggc aggaggatag cttgagccca ggaggtcaag
                                                                        240
gttgcagtga gccgagattg tgccactgca ctccagcctg ggcaacagag caagaccctg
                                                                        300
cctcaaattt aaaaaaaaa aannaaaaaa aaaagt
                                                                        336
      <210> 18
      <211> 614
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(614)
      <223> n = A,T,C or G
      <400> 18
ggtacactct tettegeett tgagtgeege tacetggetg tteagetgte teetgeeate
                                                                         60
cctgtatttg ctgccatgct cttccttttc tccatggcta cactgttgag gaccagtttc
                                                                        120
agtgaccetg gagtgattee tegggegeta ceagatgaag cagettteat agaaatggag
                                                                        180
atagaageta ccaatggtgc ggtgccccag ggccagcgac caccgcctcq tatcaagaat
                                                                        240
ttccagataa acaaccagat tgtgaaactg aaatactgtt acacatgcaa gatcttccgg
                                                                        300
cctcccgggc ctccattgca gcatctgtga caactgtgtg gagcgcttcg accatcactg
                                                                        360
cccctgggta gggaaatgtg ttggaaaaga ggaactaccg ntacttctac ctcttcatcc
                                                                        420
tttctctttt ccctccttac aaactaaggc tttngctttc aacatcgcta tgtgggccct
                                                                        480
aaaatctttg aaaattggct ttttggaana cattgaaaga aactcctgga aactggtcta
                                                                        540
gaaagneeta attgettett tacaetttgg neennenggg aetgatggga ttteanaett
                                                                        600
tcttgggact ttna
                                                                        614
      <210> 19
      <211> 296
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(296)
      \langle 223 \rangle n = A,T,C or G
      <400> 19
actititit titititit tititititgg gatggagtet cactnigitg ccaaggetgg
                                                                         60
agtgcagtgg cataatttcg gctcacttca acctctgcct cccgggttca agcaattctg
                                                                        120
cgtcagcctc cggaggagct aggactacag gcatgcacca ccatgcccaa ctaatttttg
                                                                        180
natttttagt agagatggag tttcaccata ttgaccaggc taggctggtc ttgaactcct
                                                                        240
agcctnaggt gatctgccca cctnagcccc ccaaagtacc tcggccgtga ccacgc
                                                                        296
```

```
<210> 20
      <211> 565
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(565)
      <223> n = A, T, C or G
      <400> 20
accaattata atgcattatt atgaaatatt taaaatgggg aatccaagat gacatagttt
                                                                         60
ttaactcatc cacatactgg aagtttagag aaactcagaa tttcttattt cttttcttt
                                                                        120
ttcctccata gcataaaagc tttgctaata agaataaata tatatattqq aqttttaqtq
                                                                        180
tttgatcctg tgatcagttg taaccatgtg tcataaaact ctctcacaga ttccatcttt
                                                                        240
cccaaatctt ctgatcataa cacagattgc catatagact tcccttgtaa ggagaatatg
                                                                        300
ctggccataa ggcaagcana agtgaacttg cagtttcact tcttggaaat taatgcattt
                                                                        360
gcattgactt ctataannta atctctcctg aatttttttg cttagtcaac ttactgtgtg
                                                                        420
caaagncaac agnaaattgt ctttggttna acttttaaca ggncaattta taaattgqtt
                                                                        480
tgaagaagen teeenaaatt ttttattgaa ggetgaatte aageeteent taaaatggne
                                                                        540
atngnataan gggaatttat tgtng
                                                                        565
      <210> 21
      <211> 582
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (582)
      \langle 223 \rangle n = A,T,C or G
      <400> 21
ggtactggaa caactataag acccctgttc agattaagga atttggcgca gtttcaaaag
                                                                         60
tagactttte teeteageet eeatataatt atgetgteae agetteetea agaatteaea
                                                                        120
tttatggccg atactcccaa gaacctataa aaaccttttc tcgatttaaa gacacagcat
                                                                        180
actgtgctac ttttcgacaa gatggtagat tgcttgtggc tggcagtgaa gatggtggag
                                                                        240
ttcaactttt tgatataagt gggagggctc ccctcaggca gtttgaaggc catcaaaaqc
                                                                        300
agttcataca gtagatttta cagctgacaa atatcacgtg gtctctgggg ctgatgatta
                                                                        360
tacnagttaa atttatgggg atattncaaa cttccaaaga aaattttgnc catttaaaag
                                                                        420
aacactctng antatggnga aggtgnggnt tgtgcctaac caaacttaat tccgggatct
                                                                        480
tttttatnta conggatton tttggatott nonggtaaaa aanggttgga tnocconaac
                                                                        540
nnattgaaaa nngttctntc cnnttgacct nggccanccn ng
                                                                        582
      <210> 22
      <211> 349
      <212> DNA
      <213> Homo sapiens
     <400> 22
actitititt tittititt tittitigaga tggagtetig etetiqtiqe eeaqqetqqa
                                                                         60
geaaceteeg ceteetgggt teaagtgatt eteetgeete aaceteeega qtaqetqqqa
                                                                        120
ttacaggtgc ccgccaccat gccgagctaa tttttgtatc cctagtaaag acggagtttt
                                                                        180
```

```
gecatgtigg ceaggetggt etegaactee taaetteatg atetgeteae catggeetee
                                                                           240
caaagtgctg ggattacagg cgtgagccac tgtgcccaac cctcttttcc tttttcaaat
                                                                           300
gtcaatggaa agttgattgg aaaggacaat ttggctacct tttggtacc
                                                                           349
      <210> 23
      <211> 576
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (576)
      \langle 223 \rangle n = A,T,C or G
      <400> 23
acctqttctt ggagccaatg tgactgcttt cattgaatca cagaatggac atacagaagt
                                                                            60
tttggaactt ttggataatg gtgcaggcgc tgattctttc aagaatgatg qaqtctactc
                                                                           120
caggitatitt acagcatata cagaaaatgg cagatatagc ttaaaagtic gggctcatgg
                                                                           180
aggagcaaac actgccaggc taaaattacg gcctccactg aatagagccg cgtacatacc
                                                                           240
aggetgggta gtgaacgggg aaattgaage aaaceegeea agaeetgaaa ttgatgagga
                                                                           300
tactcagacc accttggagg atttcagccg aacagcatcc ggaggtgcat ttgtggtatc acaagtccca agccttcctt gcctgaccaa tacccaccaa gtcaaatcac agaccttgat
                                                                           360
                                                                           420
gccacagttc attaggataa gattattctt acatggacag caccaggaga taattttgat
                                                                           480
gttggaaaag ttcaacgtta tatcataaga ataatgccag tattcttgac taagagacag
                                                                           540
ttttgatgat ctcttaagta aatactctga ntgccn
                                                                           576
      <210> 24
      <211> 618
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(618)
      <223> n = A, T, C \text{ or } G
      <400> 24
acttaaaata aagttaacaa ttacaacaga cccaatcaca gacaatacca gcgtagaaat
                                                                            60
attaactcca gaattatgac ttttatcagg agtaggagta ggagtaggag taggtgtagg
                                                                           120
atcaatgtca tcaggatttg cttgagggat aaacaaagtt acttgtgcaa tgttggatac
                                                                           180
ttttgatgtc aaattgcttt tatctatact tttaatggca ataaatatgt gggttgcatt
                                                                           240
ttettetgag atattitetg gtttaaatge aaagettiee ttggagttgg cetectttgg
                                                                           300
tgacagatca gtagtattta cttgaagagc atcatcaaaa ctgtctctta gatcaagaat
                                                                           360
acttgcactt attettatga tataacgttg aactttteca acatcaaaat tateteetgg
                                                                           420
tgctgtccat gtaagaataa tcttatcctc atgaactgtg gcatcaaggt ctgtgatttg
                                                                           480
acttggtggg tattggtcag caagggaagg cttgggactt gtgatccaca aatgccctcc
                                                                           540
ggatgctgtc ggctgaaatc ctccangtgg ctgagtatcc tcatcaattc aggtcttggc
                                                                           600
nggttgcttc aattnccc
                                                                           618
      <210> 25
      <211> 595
      <212> DNA
      <213> Homo sapiens
```

....

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<220> -
       <221> misc_feature
       <222> (1) ... (595)
       \langle 223 \rangle n = A,T,C or G
       <400> 25
acataccacg ctgggtagtg aacggggaaa ttgaagcaaa cccgccaaga cctgaaattg
                                                                            60
atgaggatac tcagaccacc ttggaggatt tcagccgaac agcatccgga ggtgcatttg
                                                                           120
tggtntcaca agtcccaagc cttcccttgc ctgaccaata cccaccaagt caaatcacag
                                                                           180
accttgatge cacagnteat gaggataana ttattettae atggacagea ceaggagata attttgatgt tggaaaagtt caacgntata teataagaat aagtgeaagt attettgate
                                                                           240
                                                                           300
taagagacag tintgatgat gctcttcaag taaatactac tgatctgtca ccaaaggagg
                                                                           360
ccaactccaa ngaaagcttt gcntttaaac cagaaaatat ctcagaagaa aatgcaaccc
                                                                           420
acatatttat tgccnttnaa agtatagata nagcaatttg acatcnaagt ntccacattg
                                                                           480
540
ttctanttct gaaaaaggat aatccggngt aaattttccc tggattgctg ggatg
                                                                           595
      <210> 26
      <211> 361
      <212> DNA
      <213> Homo sapiens
      <221> misc feature
      <222> (1)...(361)
      <223> n = A, T, C or G
      <400> 26
actititit tittitit tittititiga gcatattata totaattiti gaaggitgta
                                                                            60
ttttctccct tgttttaatt ttctgcanat acttttttct tttttacttt ccccaattag
                                                                           120
tttgtttctg actttcttcc tcaatctctc ctgaaccatt gtttnttttt aagatcagag cagattctta ggaactttta aaactgtatg tgggtgggat tgtcacctan agtgcttttt
                                                                           180
                                                                           240
tggagagtaa ttggatggng tgataattaa ttttatgtgt caatttgaca gggtcttggg
                                                                           300
gtgtccagtt atttggttaa acattatttc tgggtgtgcc taaaagggtg tcccgcgtac
                                                                           360
                                                                           361
      <210> 27
      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(611)
      <223> n = A,T,C or G
      <400> 27
acctgttctt ggagccaatg tgactgcttt cattgaatca cagaatggga catacagaag
                                                                            60
ttttggaact tttggataat ggtgcaggcg ctgattcttt caagaatgat ggagtctact
                                                                           120
ccaggtattt tacagcatat acagaaaatg gcagatatag cttaaaagtt cgggctcatg
                                                                           180
gaggagcaaa cactgccagg ctaaaattac ggcctccact gaatagagcc gcgtacatac
                                                                           240
caagctgggt agtgaacggg gaaattgaag caaacccgcc aagacctgaa attgatgagg
                                                                           300
```

```
atactcagac caccttggag gatttcagcc gaacagcatc ccgaggtgca tttgtggtat
                                                                       360
cacaaaqtcc caaacctttc cttgcctqac caatacccac caaqtcaaat cacaqacctt
                                                                       420
gatgccacaa gtcattagga taaaatattc ttacatggan qcccangaaa taattttgat
                                                                       480
gttngnaaag ntcaccgtnt ntataanaat aaggccagtt ttttgactaa aaaaagtttg
                                                                       540
aagagettte aagaaaneta tgatttgnee caaggggeee teeaggaagn ttgttttace
                                                                       600
caaaattttn a
                                                                       611
      <210> 28
      <211> 443
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (443)
      <223> n = A, T, C or G
      <400> 28
cgtgcccaaa gcttggcaag ttttcggctt taaccacgca caccaccacc accaccatne
                                                                        60
taaataactt actgcatcct caaagcctgt tttatgggga ttgcatggtt ttatttgaaa
                                                                       120
tcacgcctgt aatcccanca ctttgggagg ccaaggcagg cagatcacaa ggtcaggaga
                                                                       180
tegagaccaa tetggetaca eggtgaaace etgtetetat taaaaaaaaat acaaaacaat
                                                                       240
tagccaggca tggtggcagg cgcctgtagt cccanctact cgggaggctg angcaaqana
                                                                       300
atggcgtgaa acttggaggc ggagcttgca atgagccgag atcgcacttg ctgcacttna
                                                                       360
acctgggcaa caaaacgaga cttcatntct nttttnnaaa nnnaannnnn nnnnnnnnng
                                                                       420
teetttggee egaceaenet tan
                                                                       443
      <210> 29
      <211> 403
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(403)
      <223> n = A,T,C or G
ggtacttttt tttttttt ttttttttg gagtgcatat catcacccca acttcggttt
                                                                        60
tttacatttt aattigtatt gnitttaatt tattitgagg caatgictca ctatgitgee
                                                                       120
caggctggtc tcaaatgaaa acaatgctat caatcacatt cttgcatagg atatgtgtca
                                                                       180
gtaatcctcc aaaatgaaca tganaaatgg aattgtcaag tcatagatta agtgcatata
                                                                       240
acttttgaat agatagtata aattttttcc ccaaatgaga attttatatt ctcactggca
                                                                       300
acatgaaaat agccatctct ctataatctt atcaaccctc gatagtgtca ttttttaatt
                                                                       360
tataattatg agtgaaaatg gtcctgcccn ggcgggcgct cga
                                                                       403
      <210> 30
      <211> 615
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
```

```
<222> (1) . . . (615)
      <223> n = A, T, C or G
      <400> 30
60
teageaggea catagatage etgtaetttg taatattett eccaecettg agaatggaet
                                                                     120
ttgtaagate egeceetge ceacaaaaaa attteteeta actecaetge etateceaaa
                                                                     180
cctataagaa ctaatgataa tcccaccacc ctttgctgac tctcttttca aactcagcct
                                                                     240
geetgegeec aggtgattaa aaagetttat tgeteaccca aageetgttt ggtggtetet
                                                                     300
tcacacagae gegegtgaca gaaaccactt gaageeeggg egeggtgget caggeetgta
                                                                     360
atcccagcac tttgggaggc tgaggtgggt ggattacctg aggtcangag ttcgagacca
                                                                     420
gcctgaccaa catggtaaaa ccctgtctct actaaaaatc aaaaaaanta accnngggtg
                                                                     480
gtggnnggca cctgtaattc agttcttggg accttangca ngaaaatcct tgaacttgga
                                                                     540
ggcggaggtg catanttgaa acaaaccttg nctcaacctg gnaacaaaat aaaaatccgn
                                                                     600
tnaaaaaana aaaaa
                                                                     615
      <210> 31
      <211> 485
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(485)
      <223> n = A, T, C or G
      <400> 31
acgcggggat aagctacaac ataaacacat ctaggttctt gttcttagaa tacagcatga
                                                                      60
agaatttgct ttcttctttc ttcctaacat tttcatgtga gatccagaaa ggacacattg
                                                                     120
tctctggcca ttcgaagaaa gaaagaaaga aagaaaaaaa aggtatttag agacagagag
                                                                     180
agaaaaaggc tgaaatgggt tcgctgggtt ctaaaaatcc gcaaaccaaa caagcccaag
                                                                     240
ttettetttt gggaettgae teagetggga agtetaetet eetttataaa ttaaagettg
                                                                     300
ctaaggatat taccaccatc cctacaatag gtttcaatgt ggaaatgatc gagttggaaa
                                                                     360
ggaatette acteacagte tgggatgttg gaggacagga aaaaatgaga actgtttggg
                                                                     420
getgttetgt gagaaccena tnggetngtg tatgtgtgga cagteetteg geeegaacce
                                                                     480
                                                                     485
      <210> 32
      <211> 780
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(780)
      <223> n = A,T,C or G
      <400> 32
cgaggtacgc gggtgtctag accttatgtc aaaataagcc caattgtatt aaagagtatt
                                                                      60
aaattgtatt aagaataaaa acacatggcc gggcacggtg gctcacgcct gtaatcccag
                                                                     120
cacttiggga ggacgagatg ggcggattac aaggtcagga gattgagacc atcctggcta
                                                                     180
acatggtgaa accccgtctc tactaaaaat acaaaaaaa aattgtccag ccgtggtggc
                                                                     240
aggtgcctct agtcccacta ctccagagct gaggcaggag aatgatgtga acccgggagg
                                                                     300
```

```
canagettgn agtgageeng agatetegee actgeactee ggeetaggeg acagagegag
                                                                                  360
actetgtete anaaaaaat aatgantaaa aaaanaagte etgeeeggee ggegntenaa
                                                                                  420
nggcgaattt cancacatgg cngcngttac tatggatccn actcggtcca anctggcgta
                                                                                  480
atcatggcat agnttttnct gtggnaaatg gtatccgtnc aantcnccna attcaaccgg
                                                                                  540
agettaannn ntaacetggg genatnnnnn netaetteat tattgentne ntatggeget
tncattggaa ctnttgcnct gnntatnatc gcccnccngg aaagnnttnn ntgggncctt ctctgttann atctnnggct tngttgggag gntnctntna gnggntngtt tnatnggtcc ngnaaatttc agcctangnc antnagcctn ttgnttaatc tccnactnna aaaaataang
                                                                                  600
                                                                                  660
                                                                                  720
                                                                                  780
       <210> 33
        <211> 742
        <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (742)
       <223> n = A,T,C or G
       <400> 33
acataccagg ctgggtagtg aacggggaaa ttgaagcaaa cccgccaaga cctgaaattg
                                                                                   60
atgaggatac tcagaccacc ttggaggatt tcagccgaac agcatccgga ggtgcatttg
                                                                                  120
tggtatcaca agtcccaage ettecettge etgaccaata eccaecaagt caaatcacag
                                                                                  180.
accttgatgc cacagttcat gaggataaga ttattcttac atggacagca ccaggagata
                                                                                  240
attttgatgt tggaaaagtt caacgttata tcataagaat aagtgcaagt attcttgatc
                                                                                  300
taagagacag tittgatgat gctcttcaag taaatctact gatctgcacc aaaggaggcc
                                                                                  360
aacttcaagg aaagctttgc atttaaccan aaaatattta taagaaaatg cacccacata
                                                                                  420
ttataccatt aaaagttnga taaaacantt tgcctcaaaa gtttccacca tggacaagta
                                                                                  480
acttggttat cctnagcaat cttgtgcctt gattactcnn ctctattcta tcctgtnaaa gcntaatctg agtaaaattt nccctggntt gtggattggc tngtnatgta atttnttaag
                                                                                  540
                                                                                  600
netggengae enetaggnaa tnneettggg egttangnee gtngeeantt gtattngtaa
                                                                                  660
tttctngaat gtnntennen nnntacengt aagnatgggn tnggnnatnn atnttttnen
                                                                                  720
tnttnatnnn cntnnannnn tg
                                                                                  742
       <210> 34
       <211> 763
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (763)
       <223> n = A, T, C \text{ or } G
       <400> 34
ggtcaaatga ggaataatga ggaaacaaaa ccatacatac aagagggatg gcacagacct
                                                                                   60
tgtgacaaag tggtcctgaa atttctggag gggaaatgaa taagaataac cgagatagtt atgcttggag gaagaggaag atcaaggtgt cctaacctac cagaaactaa gacttatgaa
                                                                                  120
                                                                                  180
accttagtca ttaaaatatg tagtattagt tcagaaatag taaataaatc aatgtaactg
                                                                                 240
aatggaacct gggaacaaat atagctacat gtaagatctg ggtatatgct ggaggtgaca
                                                                                 300
taacaaatga agagaaacaa tggactattc aaagctgtgt tgctatcttt attggcaaca
                                                                                 360
aatatgggaa aaaatnaaat gagatcctat tcacatgaat gacaaaaata aatgccatat
                                                                                 420
tgattaaacc taaatatgac aaggaaggcc tcaaatttta gaaaaaaatg ccaaattnta
                                                                                 480
```

```
cncattggga gataattcat taacaagacc aanaaccnta aggaaagatg ntaatttnga
                                                            540
tatattaaga titactatgt tiataaatca aggatagtcc cgcttaagan actitctitt
                                                            600
atttttaatt aatattatta atatttgana cttgcttgnt tnggtgaacc ggtaatttgg
                                                            660
tattnacctt ctccggttan gattnnctaa nccntgtgnt nngttgnncc ncncnatttt
                                                            720
thtacagtth ttgcgcgnta ttncnggnng ccccennngn ngg
                                                            763
     <210> 35
     <211> 767
     <212> DNA
     <213> Homo sapiens
     <220>
     <221> misc_feature
     <222> (1) ... (767)
     <223> n = A, T, C or G
     <400> 35
60
ggaatggaat ggaatggaat ggaatggaat ggaatcaact cgattgcaat
                                                            120
180
240
300
360
                                                            420
ggaataattc naatagaatg gaatggaatg gaatggaacg gaatggaccg gatggaacca
                                                            480
attgtaatgg aatggaattg atggaatgga atggaatcac cctagtcaan ggaatgtatg
                                                            540
gaccggattc aatgaatgga tattccgnat ggatggatgg gaatgaattg atgattggat
                                                            600
ggatggatca ccatccatga agattgatga tggatgatgc cacccatgat gattatgnat
                                                            660
tagngtnata tetneatnna ggatgntnen attatgngnt gatgaeatga ntanneenne
                                                            720
netttnanen tatttttttg ggnececete ceagttgntt taaannn
                                                            767
     <210> 36
     <211> 608
     <212> DNA
     <213> Homo sapiens
     <220>
     <221> misc_feature
     <222> (1)...(608)
     <223> n = A,T,C or G
     <400> 36
acatatagtc aacgaaatat tcaaagaata actttatata ctcttgttct ttaaattcta
tectetett cagaattett ceatttaagt ttgggtattt teetagttte aacagatgaa
                                                            120
cagaagactt cattgaacat tttgacagta agctactaga gaccaattat caactggtgc
                                                            180
tacacatgct gtgttatctc ccttactatt aaactataac cctctcttgc tattttgttt
                                                            240
catgcatcac caaccaaact tcattttttc taataaaaaa taaatatata aagaagacac
                                                            300
tgacaggcat atattcacaa gatctcaact tcttaaaaca taagtatggg tatatttatt
                                                            360
tctctcaaat gcatacnaga caataattac ncagcaacca atcttttgtt caacaatgat
                                                            420
ttgantcata agcatttgga aattacataa tttcatatca atanccctgt tttttnaata
                                                            480
cagaagtaaa aaanccccaa taaccaatct taaatttcna ttatcccctt acctccaacc
                                                            540
tttnaaaggt cccaccgggc cttttccnac attaatttgg tnaaactggg gttnaaaacc
                                                            600
gcctnccn
                                                            608
```

```
<210> 37
       <211> 245
       <212> DNA
       <213> Homo sapiens
       <400> 37
acagacatgg cggcggcttt tcggaaggcg gctaagtccc ggcagcggga acacagagag
                                                                               60
cgaagccagc ctggctttcg aaaacatctg ggcctgctgg agaaaaagaa agattacaaa
                                                                              120
cttcgtgcag atgactaccg taaaaaacaa gaatacctca aagctcttcg gaagaaggct
                                                                              180
cttgaaaaaa atccagatga attctactac aaaatgactc gggttaaact ccaggatgga
                                                                              240
gtacc
                                                                              245
       <210> 38
       <211> 630
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (630)
       \langle 223 \rangle n = A,T,C or G
       <400> 38
actacactga attcacccc actgaaaaag atgagtatgc ctgccgtgtg aaccatgtga
                                                                               60
ctttgtcaca gcccaagata gttaagtggg atcgagacat gtaagcagca tcatggaggt
                                                                              120
ttgaagatgc cgcatttgga ttggatgaat tccaaattct gcttgcttgc tttttaatat
                                                                              180
tgatatgctt atacacttac actttatgca caaaatgtag ggttataata atgttaacat
                                                                              240
ggacatgate ttetttataa ttetaetttg agtgetgtet ecatgtttga tgtatetgag eaggttgete eacaggtage tetaggaggg etggeaactt anaggtgggg ageagagaat tetettatee aacateaaca tettggteag atttgaacte tteaatetet ttgcaeteaa
                                                                              300
                                                                              360
                                                                              420
agettgttna gatagtttaa geegtgeata aattnactte caaatttaca taetetgett
                                                                              480
anaaatttgg ggggaaaaat taaaaaatnt aattggccag gatnttggna atttgttata
                                                                              540
atgaatgaaa cattttngna ttaaaaatca nattacttnt aanctttgat aaantaaggc
                                                                              600
atggntgggg gtaattgggt tttttgttcc
                                                                              630
       <210> 39
      <211> 626
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (626)
      <223> n = A, T, C \text{ or } G
      <400> 39
acagtggtcc ttttcagagt tggacttcta gactcacctg ttctcactcc ctgttttaat
                                                                               60
tcaacccagc catgcaatgc caaataatag aattgctccc taccagctga acagggagga
                                                                              120
gtctgtgcag tttctgacac ttgttgttga acatggctaa atacaatggg tatcgctgag
                                                                              180
actaagttgt agaaattaac aaatgtgctg cttggttaaa atggctacac tcatctgact
                                                                              240
cattetttat tetattttag tiggttigta tetigeetaa ggtgegtagt ceaactetig
                                                                              300
gtattaccct cctaatagtc atactagtag tcatactccc tggtgtagtg tattctctaa
```

360

```
aagctttaaa tgtctgcatg cagccagcca tcaaatagtg aatggtctct ctttggctgg aattacaaaa ctcaaagaaa tgtgtcatca ggagaacatc ataacccatg aaggataaaa
                                                                          420
                                                                          480
gccccaaatg gnggtactga taataacact aatgcnttaa gatttggtca ccctctcnct
                                                                          540
aagggagccc attgagccna nggngctaaa gcctcatact ccacctgaat ggttaggaga
                                                                          600
aaatttatcc caaaaaaaaa aaaaan
                                                                          626
      <210> 40
      <211> 645
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(645)
      <223> n = A,T,C or G
      <400> 40
cgaggtacgc gggcaggaca tttaaaaggt ttcagcagaa atcttatgat tatgtctgac
                                                                           60
ttgcagtatt ttatttgcct ctttgacggc ttttttttt tttttttg agacagagtc
                                                                          120
tcacactgca ctccagcctg ggtgacagag tgagagactc cgtctcaaaa atgaatgaat
                                                                          180
gaatgaatga atgaatgaac aaacgaacaa ggtggtttaa tgtcagaaaa cttcctaagc
                                                                          240
atttgctccc caaacctttc atgtttttca agaagccttt attacataaa ggggaataga
                                                                          300
attaaaatgt ttctttataa gaaaaatata catatttgtg ttcttggccc cattaaaact
                                                                          360
aatcagtagt cetttggeca aaaaatagte aacaaganaa etgggtatga nteenggent
                                                                          420
tactcctgnt cataagtgng gatgentgtg tetganeena actgneteaa etngagetet
                                                                          480
tggggtataa caanaaaccc gngttttcat gaaacccctg ggccnttata aaaggtttcc
                                                                          540
cttggggggc ccaatgctta ttntngattn gggttccaaa anntngcaat tggnataggt
                                                                          600
gcttgaaata accccctttt agtnnaattc cnaccaaaac cntgn
                                                                          645
      <210> 41
      <211> 616
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(616)
      <223> n = A, T, C \text{ or } G
      <400> 41
acgcggggct cttcacgagg tggaaacaag atggaggatt cggcctcggc ctcgctgtct
                                                                           60
tetgeageeg etactggaac etceaceteg aetceagegg eccegacage aeggaageag
                                                                          120
ctggataaag aacaggttag aaaggcagtg gacgctctct tgacgcattg caagtccagg
                                                                          180
aaaaacaatt atgggttgct tttgaatgag aatgaaagtt tatttttaat ggtggtatta
                                                                          240
tggaaaattc caagtaaaga actgagggtc agattgacct tgcctcatag tattcgatca
                                                                          300
gattcagaag atatctgttt atttacgaag gatgaaccca attcaactcc tgaaaagaca
                                                                          360
gaacaagttt tatagaaagc ttttaaacaa gcatggaatt aaaaccggtt ctnaagatat
                                                                          420
ctcctccaac tctaaanaan gaatataaat cctatgaacc aagctcgcct tttaacagtt
                                                                          480
tgattcttcn tactgatcca aaataagcgg ttttacctcc ttattgggag acattnttta
                                                                          540
aaaaagaaag tccatntntg naaccttttt ccaaaatttn tcagananac atgctgnttg
                                                                          600
gngacggctt aaaatt
                                                                          616
```

<210> 42

```
<211> 259
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (259)
      <223> n = A, T, C \text{ or } G
      <400> 42
ngtacggtcg gtggcagtgc tattctgaga tctgtagatg cttagaatat cagtattttg
                                                                          60
gatgttgctg cattttacaa tttatttgga gtcttccttn attttcctcc agatatatga
                                                                         120
aaatatgcaa tacctgctta tatcatgtag aaaagcttag caattattaa titttctnta
                                                                         180
tttcatttta tttgaccaaa gtcggtgctt cacttgactc antgtgtttt aggtgttngt
                                                                         240
ntttntacct ttccggtca
                                                                         259
      <210> 43
      <211> 509
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(509)
<223> n = A,T,C or G
      <400> 43
acgagtgtat ttttgatggg aaggccatgc taaatctata aaacagatgt ttcctctccc
                                                                          60
aacagtggtc accagtagtt tcaacttttt ccccccagta gcatcaacca aacttagcat
                                                                         120
agtgattttt aactettige teccacaege acteatecca aetteceege tigececaet
                                                                         180
ccctgggggg aaataaccct gcctttaaaa taaatagcaa ccaagtgctc agttctatgg
                                                                         240
aaagtatgaa tatttatttc aggetttega teccaatega tttcaaaaaa caaagtetga
                                                                         300
tttctctcct cagagcagct gaggcctcca tgttacgatg gtttcatgga gattgaagga
                                                                         360
gcacatttca tcaggettag cacaaagtee etgatgeeca ceatgteeca geettagnaa
aggaaagaaa cagaattcac caccatgggg ctgaacgaat gccacaccta atgtaaatga
                                                                         420
                                                                         480
ncagctaacc ttggccaaat tgtggtttt
                                                                         509
      <210> 44
     . <211> 544
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(544)
      <223> n = A, T, C \text{ or } G
      <400> 44
ttttttaaaa gtgtcactna ntctttaann anatncatta ccattttttt tncaaantaa
                                                                          60
attacggttt taaanggaan acacatggna atntananaa ncaccgnnga annttaanta
                                                                         120
cctngggngc gancanactn anggcgaatt cgaaccaatg ggggcngnaa cnaggggatc
                                                                         180
ccagcinggt accaaaattg gcginatgat cgcaatagcg gtacctgtgn naaanggtta
                                                                         240
ttenntngta aaancagann tentnnaagn nngaccaaaa aangtaaate etggggtgee
                                                                         300
```

```
taatgannga thtaaancha ttaattgggn tgcccacctg chantttatc gttcaaaaac
                                                                        360
ccgttaaacn ngtgnaaaaa tgaatngcca accentngga aaagccgnat entttgggng
                                                                        420
cttttccttt ttggtcctna cncttcctan nngnnngttt gggnncggnt nagttcntaa
                                                                        480
aggcgnaaaa catttacaaa aataggggaa ancccgaaaa acatttaccc nagccacctt
                                                                        540
nten
                                                                        544
      <210> 45
      <211> 630
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(630)
      <223> n = A, T, C or G
      <400> 45
ggtactetet ateaetgaca aatgeagget ggattettat tatataeaga gatggeteaa
                                                                        60
aaatggggtt tcagatcttt gtgacgaaat agaatactgt ttcatatttg aatcagaggg
                                                                        120
cttcttgttc tgagaaatag gttcaaaatc attggaacca ggaacaagaa tagcttattg
                                                                       180
ttatctgtga taacactgtt ttctaaacac aaggattttc ttttttatta atatgcaaca
                                                                        240
tagacattgc cataacagaa taataaacca catgtggggt tttaaaaaatg aaatttggct
                                                                       300
aataggagca attcagctat ttttctatca agaaattggg tggggtggga tagaaagaaa
                                                                       360
aaccgggttc aaccccactt ctgcccccta accagctata tggcctggat ggagcattca
                                                                       420
acctttaata agggtcaatt tcntctgttn aaaagacccc aaacctggaa atcacnttng
                                                                       480
cctctccctg aaaataanaa ggctngattt ttggaataan aaacataatg nangctnggc
                                                                       540
ccaatggete gecegtaat ccaecetttg gaggecange ggneggaeae ttgaggtagg
                                                                       600
agttgaacca cccgccacct gggaacccnn
                                                                       630
      <210> 46
      <211> 622
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (622)
      <223> n = A, T, C or G
      <400> 46
tttttgactc ccaaagtcat tttatttaac aaaggggtca aggcagagga aagtttccct
                                                                        60
taatateeee acaactgete cacatgtett etgtggaaac actteaceag gaactagete
                                                                       120
aacactettg ctaacaattt agtgtetata caggaagget ggtgtetetg ttacaggtgg
                                                                       180
cccgttcctt aaagccttta gggttaatcg cagctgcact gagtggccaa gcagaccctg
                                                                       240
ttgggatgtg aaagcagttt gttaacaggg cccctggccg ggcccagagg ctgtcagact
                                                                       300
cancaagtaa cactgaatgt ccaaaaatac ggctgtgtta aactaacaag ccaatccttc
                                                                       360
tgctcagatc tctggataga aatgattttt cttttatcta tgggggaatg caatttcatc
                                                                       420
acaacccctt acataaacgc tcctgaaacc ctttcagtag acagcatttc aattcaaaaa
                                                                       480
ccaaaagtga aactatcttt gaaaacangg acctggctgg gaaaccatgc acacctcggc
                                                                       540
gaacactttt ccccccacg aacttggact ttntgggaag gtggcgggtt tttggcnaaa
                                                                       600
acattettga agentaggaa gg
                                                                       622
      <210> 47
```

```
<211> 253
      <212> DNA
      <213> Homo sapiens
      <400> 47
ggtactttgg tttgaaaaca acacttagag cctccagata acttttaaga cttatttagc
                                                                           60
tttgtgggtg gtattttcat gcaaataagt aagggtgggt tttatatttt gtaqaaqttt
                                                                          120
teggtectat tttaatgete tttgtatgge agtatgtata tattgtgtta agttecteaa
                                                                          180
gaatctcctt aaaaactttg aagttaatac ttttgtgcaa ctgtgttttg aataaaqcca
                                                                          240
tgacagtgtt aaa
                                                                          253
      <210> 48
      <211> 607
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(607)
      <223> n = A, T, C \text{ or } G
      <400> 48
acttacatat cctacatttg actacattat ttccaaacca agtattccat ccaaaggaac
                                                                           60
atactgctat catagagacc aaggagggac tgtttaaagt tgccaaggtg aagcgagctg
                                                                          120
agaggetttg teetegtgee agtaactetg aaatttetet taatteetge tgtegaggea
                                                                          180
gcagaatgcc atggtttccc caagtaggta gctgctttag cagttaaagc ccaaatgtct
                                                                          240
gttctgttga tcaagaggtc tctgaatttc tgaagtggtg tttcgtttct ggtgactgag
                                                                          300
ttaatccttt acaatnocto ttgtaaagtg tgctaataga aagaatccac ctttcaaagc
                                                                          360
tgcagaacca naccgtgccc taaattgacc aaccgtanct gatgtgcctn angaagtctt
                                                                          420
ttgccaactg ccctgtgaan accectnett cccccaget ngtggettge acactgaaca
                                                                          480
tttaaactgn gcaaagccgt gtagttataa nacagtaaat cccaaggctt ggttaantgc
                                                                          540
tgggnnaaaa ctggttggat anacttaact taaaacccct tacataaacn tnggaactcn
                                                                          600
aagaaaa
                                                                          607
      <210> 49
      <211> 421
      <212> DNA
      <213> Homo sapiens
      <400> 49
ggtaccactg gatgagggc cgggacatac tgactgcccc tttgacccca caagaatcta
                                                                           60
tgatacagec ttggetetet ggatecette tttgeteatg tetgeagggg aggetgetet
                                                                          120
atctggttac tgctgtgtgg ctgcactcac tctacgtgga gttgggccct gcaggaagga
                                                                          180
cggacttcag gggcagctag aggaaatgac agagcttgaa tctcctaaat gtaaaaggca
                                                                          240
ggaaaatgag cagctactgg atcaaaatca agaaatccgg gcatcacaga gaagttgggt ttaggacagg tgctgttccc gagactcagt cctaaagggt ttttttccca ctaagcaagg
                                                                          300
                                                                          360
ggccctgacc tcgggatgag ataacaaatt gtaataaaag taacttctct tttctttcaa
                                                                          420
                                                                          421
      <210> 50
      <211> 624
      <212> DNA
      <213> Homo sapiens
```

```
<220>
      <221> misc feature
      <222> (1)...(624)
      \langle 223 \rangle n = A,T,C or G
      <400> 50
ggtacttcag tattgcattc tattcctctt aatgttttta tgggatctcc agggaaagag
                                                                          60
gaaaatgaaa accgtgatct aacagctgag tctaagaaaa tatatatggg aaaacaggaa
                                                                         120
tctaaagact ccttcaaaca gttagcaaag ttggtcacat ctggtgctga aaqtggaaat
                                                                         180
ctaaatacct ctccatcatc taaccaaaca agaaattctg agaaatttga aaagccagag
                                                                         240
aatgaaattg aagcccagtt gatatgtgaa cccccaatca atggatcctc aactccaaat
                                                                         300
ccaaagatag catcttctgt cactgctgga gttgccagtt cactctcaga aaaaatagcc
                                                                         360
qacaqcattq qaaataaccq qcaaaatqca ccattqactt ccattcaaat tcqtttattc
                                                                         420
aaacatgatc aagaaacgtt ggatgacttt aaaaaanatg ccntaaggac anttgtqatt
                                                                         480
tgcaggtggg aagatnaaca gttcatatcc actgaatgaa atgcatcttg tggaaganct
                                                                         540
catgnatnaa ggttaatggc tgaaatgaaa actccaaaag aaaccaaaaa ataccggccc
                                                                         600
ctttgaaatt caggganncc tatg
                                                                         624
      <210> 51
      <211> 632
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(632)
      <223> n = A, T, C \text{ or } G
      <400> 51
ggtacgcggg ggaaacggaa gtgagcggcg gggtcgactg acggtaacgg ggcagagagg
                                                                         60
ctgttcgcag agctgcggaa gatgaatqcc agaggacttq gatctgagct aaaggacagt
                                                                         120
attocagtta ctgaacttte ageaagtgga cettttgaaa gteatgatet tetteggaaa
                                                                         180
ggtttttctt gtgtgaaaaa tgaacttttg cctagtcatc cccttgaatt atcaagaaaa
                                                                         240
aaatttccag ctcaaccnaa gataaaatga attttttccc cctgaagaaa cattcagggc
                                                                         300
tattttgctt cccttaaaat accagaatgg gattcaaggg cagtgccacc aggtcaaccg
                                                                         360
ettteattte ttteaageet caaatettte acttgaatgt ttgaagggta atggatgaag
                                                                         420
acctattgga attgagggat atctttaatg atccgccca aaccgaatcc ttggaaaagc
                                                                         480
caccettgat ggtggaatat aacettggtt actgaatatg tgcctgtcat ggaaccgagg
                                                                         540
cegeatetgg ttatageate tttgacetge eggeegeece aaaggegaat ceaeneetge
                                                                         600
ggccgttcta tggaccaact cggnccaact qn
                                                                         632
      <210> 52
      <211> 623
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (623)
      <223> n = A, T, C \text{ or } G
      <400> 52
```

```
acttttaatg gtgggaattt acagtagaag catcetttge tgagttatae atteetttat
                                                                              60
caatctcttt tgatacaaca tttaaaacaa gtagcttcaa gaaaccactg gtgttttgag
                                                                             120
gatagtattt ctaaatagca ttcaggaaca gagtattatt gcacagatct gaagatcaaa
                                                                             180
aaaaagetea aggaaataca gateggaagt getgatgagt tatatttatt gaaaacceaa
                                                                             240
cttttaagga agtgctaaga tcagtcaccc atgtgaataa gaagccagga aaggaaagat
                                                                             300
ggggaaagcc canatcacca ggcttctatt aaggaggaaa gcaacagang aaacagtqaa
                                                                             360
agggaacaga aaggggtagc caagtgttac aaaaaanccg actggataac caaactncaa
                                                                             420
aaagngtatg ttggggagaa ctgaaangga aaacaaaata cttgactaat cntaagtaga
                                                                             480
aaaaagcagn tagagaaaac caaatatttc tggncctgtc acatacaact tcaaataccc
                                                                             540
ttatanaatc caaaaatgat gtgtgtaagg naaaatttat tgccntccga aaaataantt
                                                                             600
tntccaatnt qaaacaaatc aac
                                                                             623
       <210> 53
       <211> 627
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(627)
       \langle 223 \rangle n = A,T,C or G
       <400> 53
ggtacgcggg gtegcatgcg ctgtggctaa tgccgtaggc tctttcaggg ctgagccatc
                                                                              60
ctgcgtgtct tgcgctcggt ggaaatgccc agccgaggga cgcgaccaga ggacagctct
                                                                             120
gtgctgatcc ccaccgacaa ttcgacccca cacaaggagg atctaagcag caagattaaa
                                                                             180
gaacaaaaa ttgtggtgga tgaactttct aaccttaaga agaataggaa agtatatagg
                                                                             240
caacaacaga acagcaatat attetttett geagacegaa eagaaatget gtetgagage aagaatatat tggatgaact gaaaaaagaa taccaagaaa tagaaaaactt agacaagace
                                                                             300
                                                                             360
aaaatcaaga aatagtcaac ctgatttcac ataacaatgt gtggcatttg ttgttctgta
                                                                             420
aacttttetg ctgagcattt cagtcaagat ttaaaagagg acttactata taatcttaaa
                                                                             480
cagcggggac ccaatagtag taaacaattg gtaaagtctg atgttaacta ccagtgntta
                                                                             540
ttttctgntc acgtnctaca cttgangggg gtttgactac ccancctgtg gaagaagaaa
                                                                             600
gaagcaatgn ggttctatgg atggaga
                                                                             627
       <210> 54
       <211> 565
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(565)
       <223> n = A, T, C \text{ or } G
       <400> 54
ttttccttga gtgctccctt ttatgtcatt ttattttctt ttatgcagac cagtgggggg
                                                                              60
aaaatcccat agattcttct ggaaactgtc aagatgctgg gaagatgaat gcaaaactta
                                                                             120
catagattgg gatgtccaca gtttggattt tcaaggtatg gcttttgcag gatgacgtga tcaacccaaa cttctgcttg atctggtttg tcctgaactc ctgccacttg ccgccaacca
                                                                             180
                                                                             240
gggcctctgc tctgatctca tacttcacca ggcgtgccgn tcgcaggctg acgtggttgt
                                                                             300
getegtagae egeagagga gattecaggt etgtgtgett tattetetge atgtaaaac
                                                                             360
tataagaggt agtatcatgt ttgagtccct ttatcttaaa gaagaatcca tatagagcaa
                                                                             420
```

```
tcgttttcga ataagttgna ttctctgngt ctggcactgt gtccagtgct ctcanaggat
                                                                        480
gcangggaga anaccaaaaa gtntctgagc agtctcacat gggaaataaa atgtgtcccc
                                                                        540
ggtaccttgg ccgngaacac nctaa
                                                                        565
      <210> 55
      <211> 451
      <212> DNA
      <213> Homo sapiens
      <400> 55
acagagatga caagagaaag gcacaaatga ccggagtcag ggattgtggt gagggctcca
                                                                        60
catgaagaca gcatgttgga ggagaccaag ttgggaaggg tgacatgtca tacatcaaaa
                                                                       120
gttgccccaa gatagcaggt tataatgggc tagagagaaa ttagagggaa catctcttcc
                                                                       180
ttcacttgaa caacaccaaa aatagaagac cagagaatag aaggatggtg acaaatccca
                                                                       240
aaaaggaaat ggaggaggag ttcgtggaag ggcagaaaca ctttaatcct agagggaggg
                                                                       300
tgaggcactg ttgaaaagag aagcaaactt tggcaggggt ggccattctg ccttgctgag
                                                                       360
teatgggetg agatacggaa gteactttea atcattttet actteteeca gggeacteag
                                                                       420
acaaaatcag tgcaaggtat atggaagtac c
                                                                       451
      <210> 56
      <211> 623
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (623)
      <223> n = A, T, C or G
      <400> 56
ggtacgcggg gcttccgaga cgcactgggg gccggatgta gaatcctgct tatctgtgaa
                                                                        60
atgeagttaa cacateaget ggacetattt ceegaatgea gggtaaceet tetgttattt
                                                                       120
aaagatgtaa aaaatgcggg agacttgaga agaaaggcca tggaaggcac catcgatgga
                                                                       180
tcactgataa atcctacagt gtttcactct tgttgcccag gctggagtgc aatggcgcga
                                                                       240
tettggetea eggeaacete tgeeteeegg gtteaageaa ttgteetgee teageeteet
                                                                       300
gagttgctgg gattacagat tgttgatcca tttcagatac ttgtggcagc aaacaaaqca
                                                                       360
gttcacctct acaaactggg aaaaatgaag acaagaactc tatctactga aattattttc
                                                                       420
aacctttccc caaataacaa tatttcagag ctttgaaaaa atttggtatc tcaacaaatg
                                                                       480
acacttcaat tctaantgnt tacattgaan aagggagaaa acnataaatc angaatacct
                                                                       540
aatatettea gingaangge aleaaggite teitgaaaac itneeggaat aatgaainin
                                                                       600
ccnaagtcca aaanattttt aac
                                                                       623
      <210> 57
      <211> 622
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(622)
      <223> n = A,T,C or G
      <400> 57
```

```
cgaggtactt ttttttttg tttttttt tttggtttct gtcctttaat tttttaacag
                                                                        60
aatatacaga gccacacaat acgatttcaa tttcaaatta tgggagatca tattcaaata
                                                                       120
tgcttaggtt tgacaagttg ctgttacaat actqaqaact ttcatqaaaa cqqtatttaa
                                                                       180
caatttttaa gataatcaaa tatctttttg ctacgtgggc caacgcatta atactaactt
                                                                       240
gtttaaaaaat gcagtctttt agacttcaaa ttattataaa acaatatcaa gatcatatag
                                                                       300
atatacttcc tgattactca aaactcgttc cattctgatg gaggctgaag gtaaatgtta
ttatacatta gaacatttca tgaaaccact tctcctttgc acttacctgt aaaagtcaaa
                                                                       360
                                                                       420
aattaaacca caatttccta agacataact atttctagaa tacattggtg taatcataaa
                                                                       480
540
aggaccnaaa aaaattggga atttggattc cttacataac aggactcata cttctgattt
                                                                       600
aataaattnc actctttca ag
                                                                       622
      <210> 58
      <211> 471
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (471)
      <223> n = A,T,C or G
      <400> 58
ggtacttttt ttttgtttgt tttctagact taataaaagc ttaggattaa ttagaagaag
                                                                        60
caatctagtt aaatttccca tttgtatttt attttcttga atactttttt catagttatt
                                                                       120
tgtttaaaaa gatttaaaaa tcattgcact ttggtcagaa aaataataaa tatatcttat
                                                                       180
aaatgtttga ttcccttcct tgctattttt attcagtaga tttttgtttg gcatcatgtt
                                                                       240
gaagcacccg aaagataaat gatttttaaa aggctataga gtccaaagga atattctttt
                                                                       300
acaccaatto ttootttaaa aatototgag gaatttgttt togoottact tttttttott
                                                                       360
ctgtcacaat gctaagtggt atccgaggtt cttaatatga gatttaaaat cttaaaatgn
                                                                       420
ttcttatttt cagcacttac atcatttggt acctgccngg cggccgntcg a
                                                                       471
      <210> 59
      <211> 618
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (618)
      <223> n = A, T, C \text{ or } G
      <400> 59
ggtacataca caatcactca actggaacaa tcaaaaccat ctatgagtgt ggttattaaa
                                                                        60
aaataaaatt acgttcatac aatggtagaa aatgaaatgt ttttattaat ttgattatta
                                                                       120
atacaaaacc acacatatat gaattatata acctagtgtt atatatttaa aaatctttat
                                                                       180
gcttgcaact gaaatgtctc tactccaagg gaagtttctg atttttaatt ttcttatttt
                                                                       240
aaggaateta ttatatteae aatgattaaa atgeettaca cataggeaaa aageagaeee
                                                                       300
aatcccagca aacagaaaaa ccataagtct atcatatcac catatgtttc accatatagt
                                                                       360
tttgaaaaat aatcctattt gcagtttggt atgtcttcat atttatactt attatcaaag
                                                                       420
tgattgcata ttgaggcaca gagcttaaag aggaaatata tattacttat aggggaacca
                                                                       480
gacactgaaa caaggaatat caatcaatgg cttcaaacna aaaaaaaann nnnnnnnnn
                                                                       540
nnnnnnggaa aaggaaaagt cctgncccgg cggncgttca aagggcnaat tcaaccactg
                                                                       600
```

```
ggggccgtac ttatggac
                                                                               618
       <210> 60
       <211> 606
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(606)
<223> n = A,T,C or G
       <400> 60
actitttaaa ccctcccaac cagccctttc tcaatattca tcaaatctaa aacatttagg
                                                                                60
gggcaaaatt ctaacatgtt catggtatct tgcaaatagt aaaagcttta ttctgaagga
                                                                               120
ttataaacta gttttctcca ttttaactag cactatttig tggaaattag aaacctcttt
                                                                               180
tatttctctt cccaaaagta atacttatta taaggctgta gtatcaggtt aaggatacag
                                                                               240
ataaataaag ttcacttata tcttcttaca aatgtctggg ttttaatatg gttaatcact
                                                                               300
tatatacaaa tattacaact ttttagtgca agtttttgga agaaaacttt ttgataaaac
                                                                               360
actgtgattg atgtgacttt atttttaatt taaacgatga ggtggccaga agaaagatgg
                                                                               420
gtctaaaatt tctcccatga aagatgtaaa actatggctt ttttaaaatc aaaatttcat
                                                                               480
ctttaaaata atgggttgaa atctggatng gatctgaaca gaataatcac atttaggatc
                                                                              540
tatataaatc tcaactggag tntaactgaa ggaaataccn ngattttaag aaatatnttc
                                                                               600
aaaaan
                                                                               606
      <210> 61
      <211> 620
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(620)
      <223> n = A, T, C \text{ or } G
      <400> 61
ggtacattct ggtatgaaaa catctcaaaa tgtaacaaca caagagtttg ggtcaagacg
                                                                               60
acccacccag gaggetgtaa aaactggttt gaactagaac tgtggaatgg aactagttta
                                                                              120
aaatatgaag cagctctaaa caccaagctt agagacattt gccctattag aaaacaaaaa
                                                                              180
tcattaaagc tacaaaataa caagtgcaaa catgctgaac ctgtttccag ggagtgacat tcccttctgc caacaggtcc caaactcaca cccacaaggt gtaactctct ttcctgttccactagatttc ctttctctca tctcaaaggt cctcagaaat gacaatggaa aacgtatgaa
                                                                              240
                                                                              300
                                                                              360
ttgttgaaat ttaccetgtg gaccaattce tgaagagata acagccacaa etetgagatg
                                                                              420
attaagacat gcagtgttta cttgatgact ttctgnattt ctagaaaccc tcaaagcatt
                                                                              480
aaactgncta tttcaaaatc taaacttnct agcactttta ttatttggag taagcnnacc
                                                                              540
gaagacaatt tactggccca caggaataac cacgcttact tgtcaccata agtttacggn
                                                                              600
atggacattc actggaaaac
                                                                              620
      <210> 62
      <211> 614
      <212> DNA
      <213> Homo sapiens
```

```
<220>
       <221> misc_feature
       <222> (1) ... (614)
       <223> n = A, T, C \text{ or } G
       <400> 62
 geogaggtac ataaatetgt gateceattt ettattgeac catteaggaa caetttatat
                                                                         60
 aaatgagtgg ctttttattt catattatta gtagtatcat ggttccatta caggcctatt
                                                                        120
 aacatcatac attgtcatta gtctttgaag aaaaaatatg taaatatata tgtgtaacat
                                                                        180
gagaatttet etetaaagea gggettaaaa ttttttggaa aagtttgaca aageatacea
catgaattca gatttacctc aatgctaaga attatgtta gttaggaaaa aggaaagtca
                                                                        240
                                                                        300
ttttgacctc aggtagaaaa atagattgct ttgagtttta tgtagcttta gactttaaaa
                                                                        360
agttagaatt tattctgtaa ctaaaaatta tttgaaaaaa ttatgcctct ggtttaatta
                                                                        420
ttggtgatta cacactettt etettaceet tgngtattga actatgteca taatcaagtt
gatgtggatc ctgaaaaatg gtatgaacat ctgatgggat tggcacatta ttttaaaant
                                                                        480
                                                                        540
agcatctgac acttcaaaac tgtcantgng atgggttcac cataccacgg ntgaccntac
                                                                        600
attaaatttt nacn
                                                                        614
       <210> 63
       <211> 616
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (616)
      <223> n = A,T,C or G
      <400> 63
ggtacatata agagtaatta gttttattct ctctttttta taaaatcggg tttcagatga
                                                                         60
gatgtttatc ttagactatt ttagggaaaa attttacatg tttgagatgg tggagtaaaa
                                                                        120
agactgttaa acatttcttt taaaaaatta tttttacatt acaacaatat atttatgatg
                                                                        180
tgttcagatc aaaaatttaa cttctgtgtc ccagatctac tttcaaagtg agattttcac
                                                                        240
tigtcagctt aaatttctga ctagaactaa catitgtgta tttttgtgct tagtcggaat
acaaatttca cagtggattt ttgaagtttg tccttaaatt ggataaaatc aagtgattaa
                                                                        300
                                                                        360
agttactaaa gagataaaaa tggtaatttc catttttaaa agtaatttgg ttgtgtttat
                                                                        420
agttatttgt acttcgagtc tecettcacc atttccgacg gcatctacng ctcaacattt
titggtaccc cangettica cggacttcac gtcattattg gctcaactti cctcactate
                                                                        480
                                                                        540
tacticatce gecactaata titeetttac atceaacate etttgaettt naageegeeg
                                                                        600
ctgatnctgc attttn
                                                                        616
      <210> 64
      <211> 612
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (612)
      <223> n = A,T,C or G
      <400> 64
ggtacagata tcattncttg tgtatgccat gacttgaaaa agtttgggaa gctctttanc
```

```
aatatcagct aanaggatat gaaatcacag gtgatagcag ttgtcattca gtaatttcct
                                                                          120
 acaagcagca ccccaaagga aatatagtcc taatctttac tatccacttc taaatttaat
                                                                          180
 gtgaatttca tacatgttat tagttgtttc ctttataatt ttataaaaat tattcatcgg
                                                                          240
 gagtttaact tecaetteea tgetategga tgtgttggge tecatgeaag aacttggaag
                                                                          300
 aaaaacaggc aggaatgcat ttgcataatg acccagatca tcattttctg caactgagaa
                                                                          360
 ttatatttca tcattgcttc tagaagtctg caattcttta cttttctttg gtgcattatt
                                                                          420
 atctangtgc ccatcactgg ataatgtgga gtgactagag aagtcatnta tcactggaag
                                                                          480
gncctgccon ggcggccgtt caaaaggnca antccagcan nctggcggcc gttctaatgg
                                                                          540
gntccaactt ngggnccaan cttggngnan tcatggcnta acnngttccn ggggggaaat
                                                                          600
gntntccctc ac
                                                                          612
       <210> 65
       <211> 599
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(599)
       <223> n = A, T, C or G
       <400> 65
acaagctaca aaatagcatc tettteatgg tatgtttgag tgtgtaattt tagtttettt
                                                                           60
tctggttgta tttgtggtag tcagatgtgt tggattgatt ccaactggac agagtaagga
                                                                          120
attccagcat cetetteetg ettgetegtg ttaccccaca gatcaaacce tcaattctag
                                                                          180
ttggggatgc tgtctagccc cacaccatga ctgaagcctt aagcactgtt gcgcctccat
                                                                          240
gtgctttggg tcagcaaccc cagtggtatt ctaccagagc attgtgggaa ggcagatgta
                                                                          300
tagtcaggtc ccaacagcaa attgttgggt gtgagagttc taaagtatag gggtgaaggg
                                                                          360
aaagagaang atatgaactc ctctgacctt aaccacattc atttaacttt tatgcctact
                                                                          420
taacaagaga acctggagaa aactatcgna ttcaagagat taatcaaaat cagggtttan
                                                                          480
ccagccatga ccgaaancne etteettaac eteatettgn anggetgnaa naatteanne
                                                                          540
ctaggatggt taanccagaa ccccngatga ttaantgtcc aaccttnatt tncatantn
                                                                          599
       <210> 66
       <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(611)
      <223> n = A,T,C \text{ or } G
      <400> 66
ncatgacett tagtggaaga ttatttggte atcaaatace catatecaag tttecatggg
                                                                           60
gcctgggaat ttcctttcac ttggatagaa agtatatatt aggaaagtcc agttaataag
                                                                          120
tattttatt taaaaaaaaa aaaaaaggaa aaaagaatca gcagaagtca agttgtctta
                                                                          180
agtettaagg etttetggat ttetteettg gaggaggtea ggatetteee aaggeetggg
teetegaata ttetteeagt cateaaactt ggagtetttg attteteat atteegaete
                                                                          240
                                                                          300
taaagatatt ttattetett teagttttt tteaagetea ggateeattt taetetteae
                                                                         360
agcatcatat cggatttgag aaaactcacg aagaccaaaa gaaccttcaa caatcagcaa
                                                                         420
caacatgggg actccatacc cagagtettg gtettgegaa aagcaegent naacegeggg
                                                                          480
tgccaacatg agtgaactct ttcatcggtt naaactccaa cnggcctacg caaactccca
                                                                         540
```

```
atttacaggt tangctttta ccaaacaagt ncctnggcgg gacnccctag gggaattcgc
                                                                              600
cactgggggg t
                                                                              611
       <210> 67
       <211> 639
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (639)
       \langle 223 \rangle n = A,T,C or G
       <400> 67
nagaattegn gettnenage ggtegneegg geaggtacae tttaettaaa aactattaae
                                                                               60
agtttttcat gttgcactgg tggtaatttt gaacttggaa ttactgggtg ggaattccag
                                                                              120
gaaccacaga gtattgattt ttgctgccaa aatgctcttg aagcagatgt ccctgtgctc
                                                                              180
cectggetge ttetggetga aggggggagg tgtagactga agettgggea etcatgtgtg
                                                                              240
tececteeca greeceatee tagragagee agreteatra gageageeata garaageerg
                                                                              300
gaacttggct gcattagtga cttgatcctg gtatgaaatg catactgggt ataaagntgc
                                                                              360
tcaagnattt tatttccttg gccacaactt ccatagatgc caatggtttg atagcctcag
                                                                              420
tttctnaacg atgtcttttg gttacagtgc tcacttantg ngagtcaaga aatgcttgag ttaccagaaa cttcttantc aggttgagta acnttttacn ttcatgngta nctnnggcgc
                                                                              480
                                                                              540
gaacaccctt anggggaatt ccacacactt ggnggccgta ctaanggatc caacttgggn
                                                                              600
ccaacttggg ggaaaaangg cnaantggtt ccttgngaa
                                                                              639
       <210> 68
       <211> 611
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (611)
      <223> n = A, T, C \text{ or } G
      <400> 68
tegaceggee geeeggeeng gneetteece ateaetnnac tggnacnate aaaacentet
                                                                               60
atgantgngg gtattaaaaa ataaaattac gttcatacna tggtagaaaa tgaaatgntt
                                                                              120
ttattaattt gattattaat acaaaaccac acatatatga attatataac ctagtgntat
                                                                              180
atatttaaaa atetttatge ttgcaactga aatgteteta etecaaggga agtttetgat
                                                                              240
ttttaatttt cttattttaa ggaatctatt atattcacaa tgattaaaat gccttacaca taggcnaaaa gcagacccaa tcccagcaaa cagaaaaacc ntaagtctat catatcacca
                                                                              300
                                                                              360
tatgtttcac cntatagttt tgaaaaataa tootatttgc agtttggnat gnottcatat
                                                                              420
ttatacttat tatccaagtg atgentattg angneenaag etttaagang gaattttntt
                                                                              480
cctatngggg accenaceet tgaccegaat teatcaangg ntttaaceca aaaaaaann
                                                                              540
aaaaaaaaat ggnaangggg ctcccttnaa anccccccca acctntttnt ttaacnagnc
                                                                              600
tnagcctttc a
                                                                              611
      <210> 69
      <211> 606
      <212> DNA
      <213> Homo sapiens
```

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<220>
      <221> misc_feature
      <222> (1) ... (606)
      <223> n = A, T, C \text{ or } G
      <400> 69
accaaagcat tacccgcatg gtagagaaca cactcgatta aaaatgttaa gctatctgaa
                                                                          60
aaataaaatg tgcaagtett caggatggca caaaacaaag gtcaatgett ettggggcac atttettaga gggettgetg agtgtgtaaa tataategae ttttgtttgt gttacatgae
                                                                         120
                                                                         180
ttctgtgact tcattgaaaa tctgcacaat tcagtttcag ctctggatta cttcagttga
                                                                         240 -
cctttgtgaa ggtttttatc tgtgtagaat gggtgtttga cttgttttaa cctattaaat
                                                                         300
360
tcacattaaa tgggtttggt ttggcttctt ttaatcaggc tttctgaaca ttgagatatc
                                                                         420
ctgaacttag agetetteaa teetaagaat tteatgaaaa qnetntnact ttgaacegaa
                                                                         480
accanaatac ctcggccgga caccctaagg cgaattccag ccactggcng gccgtactaa
                                                                         540
nggatccanc ttggtnccaa cttggggnaa catggcnaac tggttccggg gaaatggatc
                                                                         600
cccncn
                                                                         606
      <210> 70
      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (611)
      \langle 223 \rangle n = A,T,C or G
      <400> 70
ncgtggncgc ggccgaggtn ctttttttt tttttttt tttttttt tttttttnn aaaanqqqta
                                                                          60
accttaaagg tttantggcc ccccaaangn aacctggggt taatggcttc nnattttaaa
                                                                         120
tttttggaaa ttaaaaaaat tacnagtttt aaatagccna tggctggnta tgttttcana
                                                                         180
aaacatgatt agactaattc attaatgggg gcttcaagct titccttatt qqctccanaa
                                                                         240
aattcacccn cettttgncc cttcttaaaa aactggaatg ttggcatgca tttgacttca
                                                                         300
cactetgaag caacateetg acagteatee neathtactt caaggaatat cegttggaat
                                                                         360
acttttcana aagggaatga aagaaaggct tgatcatttt gcaagggccc caccacgtgg
                                                                         420
gegganaaat caettetaca ggttattace tgganngtea aagntttetg naaaacanet
                                                                         480
tgctctcaac tggtttacca tttggtgctg gagctnacaa ccggtttaag gcccttggna
                                                                         540
anggtccaag neccaanaaa ettteeeggt cetteeggng geettnaagg gaateeneee
                                                                         600
tgggggcgtt t
                                                                         611
      <210> 71
      <211> 588
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (588)
      <223> n = A, T, C or G
      <400> 71
```

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سؤلط السرز بالسد
 nctgggaacn ccgaaggtgg aaggccnttt cataacattt cttgtggatc aaaccaccgg
                                                                              60
 gacacetttt ttnccatcaa caggactage gtettgtcag tettggtgac agtgacattg
                                                                             120
 aangtggggg cccaccggtg ctcttggtac tttcccaaga ggtcctcatc ctgagacggt
                                                                             180
 ctctacccat gtttaaccca aagagtgcag gccaggttcc ttatccttct gatgaaggat gagagaactc atttagaagt cagagcaaac tagggtctca gtattgagaa acgcacctgc
                                                                             240
                                                                             300
 canggaatca cagagacatc ggggtgcccg cgatggcctc atgaaccatg cctngacggn attcaggaac cctgcaaacg tgctttttga ctcattggnc agtgtgaatt ttacacaagg
                                                                             360
                                                                             420
 naaacetggt cnaaggcatt ngggaattgc tecaacnnat acttectntt aggaacecaa
                                                                             480
 ggaancaggt tenegaattt tgaaaactgg gtntgaagtt etttetteet ttgggnacaa
                                                                             540
 ggccttaaca aanancttgn ggnttccaaa tggncctggc cccacacc
                                                                             588
       <210> 72
       <211> 591
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(591)
       <223> n = A, T, C or G
       <400> 72
ggtacaaact tagaagaaa ttggaagata gaaacaagat agaaaatgaa aatattgtca
                                                                              60
agagtttcag atagaaaatg aaaaacaagc taagacaagt attggagaag tatagaagat
agaaaaatat aaagccaaaa attggataaa atagcactga aaaaatgagg aaattattgg
                                                                             120
taaccaattt attttaaaag cccatcaatt taatttctgg tggtgcagaa gttagaaggt
                                                                             180
aaagettgag aagatgaggg tgtttaegta gaccagaacc aatttagaag aataettgaa
                                                                             240
                                                                             300
gctagaaggg gaagttggtt aaaaatcaca tcaaaaagct actaaaagga ctggtgtaat
ttaaaaaaaa ctaaggcaga aggettttgg aagagttaga agaatttgga aggeettaaa
                                                                             360
tatagtaget tagtttgaaa aatgtgaagg aetttegtaa eggaagtaat teaagateaa
                                                                             420
gagtaattac ccacttaatg gttttgcctt ngacttttgg gttaagaata tttttaaatc
                                                                             480
                                                                             540
ctgnggctnc cttaattggc cgnttgncca ngggttccnn aaatgggttc n
                                                                             591
       <210> 73
       <211> 581
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(581)
       <223> n = A, T, C \text{ or } G
       <400> 73
acgegggtat etgtaatttt tataatteat eaattetgga atgetatata taatatttaa
aagacttttt aaatgtgttt aatttcatca tcgtaaaaag ggatcatctc agagagaaca
                                                                             60
gcagtattet gegtattett aaaaatgete tagagtaaca titgaagtaa ticacigtag
                                                                            120
tgtatgccag tcctagaaat aatttttta atttctggtg tctgtttcta atacactaac
                                                                            180
caagttttca aaatatattt acaaagatgc atctttaccc attattttaa aatgattaag
                                                                            240
gaggatagtt gcttcaggta acaagctaat ttttcaaata ttaggccctt acagaactat
                                                                            300
ttagtcaaaa agtaagatat tootttaaaa tatataacco aaagotttoa gttaaaccat
                                                                            360
gatatatcac aaatactatt aaaatggtaa agagaaaatg caattgcant taatgatgcc
                                                                            420
caaatngtaa aatatngaga ttcaaaagct gggnctttat ttaggnggga tnccaatgnn
                                                                            480
                                                                            540
```

```
aatgatactg gcctggnttt acctttacct tttaaaaaan a
                                                                           581
       <210> 74
       <211> 599
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(599)
       <223> n = A,T,C or G
       <400> 74
cgaggtactt tttccgcaca tgccttgtgc ctatctgagt attgatgcca tggatgtggc
                                                                           60
cggagaacag cagctggatg tggaacacaa cctgttcaag caacgactag ataaagatgg
                                                                           120
cateceegtg ageteagagg etgageggea tgagettggg aaagtegagg tgaeggtgtt
                                                                           180
tgaccetgac tecetggace etgategetg tgagagetge tatggtgetg aggeagaaga
                                                                           240
tatcaagtgc tgtaacacct gtgaagatgt gcgggaggca tatcgcccgt anaagctggg
                                                                          300
cetteaagaa cecagatact attgageagt geeggegaag agggetteag ceagaagatg
                                                                          360
caggaaccag aagaatgaag ctgccangtg tatggctttc ttggaaagtc aaataaggtg
                                                                          420
gcccgaaact ttcactttgc ccttggggaa ganctttcca gcantcccat gtcacntcat
                                                                          480
tgacttggca aactttggnc ttgacaaccn tnaccatgac ccactacatc ancacctgtc
                                                                          540
atttngggga ggactttcna gccttgggaa acccctngac cccccaatgg taattggcc
                                                                          599
      <210> 75
      <211> 594
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (594)
      <223> n = A, T, C or G
      <400> 75
acatcaaatt ataaatgcaa aacaggttca gatttcatct tttgtgattt cttttaaata
                                                                           60
ctattcattt ttatttaaat gcacagtatt tcccctatat tttagtcctt ccattcctag
                                                                          120
agacaaacca gttatttggt ggtgggaagt agctgaagca aagaaggaaa agtaatacct
                                                                          180
ttaacctcac tagcttcaag agtagacatt cttactagct caatttaaat aattgatttt aaataggaag aaaagaggat atatttaaga tacatagaaa ttatgatgtg aagtattcat
                                                                          240
                                                                          300
gagaatetgt agattecate aaaataagta ggaacteate taaaattgtt ggatttaaag
                                                                          360
aggcactttt ggttatgatt caaatatggg gaatttgaga aatattcatt ttgnccactg
                                                                          420
gatggtcact attttactaa aanggnagct ttttatgggg ggactgngac tgaggtctta
                                                                          480
aagactgaaa gaagttgggg ggttcatttt cngtaccacc ttcnnggacc atttggacct
                                                                          540
ttggccggga acacccctaa ggngnaattn cngnccctgg gggccgtcta atgg
                                                                          594
      <210> 76
      <211> 585
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
```

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<222> (1) ... (585)
       <223> n = A, T, C \text{ or } G
acgcgggggg cggagtagca agtggccatg gggagcctca gcggtctgcg cctggtagca ggaagctgtt ttaggttatg tgaaagagat gtttcctcat ctctaaggct taccagaagc
                                                                            60
                                                                           120
tetgatttga agagaataaa tggattttge acaaaaccac aggaaagtee eggageteea
                                                                           180
tecegeactt acaacagagt geetttacac aaacetaegg attggeagaa aaagateete
                                                                           240
atatggtcag gtcgcttcaa aaaggaagat gaaatcccag agactgtctc gttggagatg
                                                                           300
cttgatgctg caaagaacaa gatgcgagtg aagatcagct atctaatgat tgccctgacq
                                                                           360
gtggtaggat gcatcttcat ggttattgag ggcaagaagg ctgcccaaga cacgagactt
                                                                           420
ttaccaaget tgaacttana aaagaaaget egtettgaaa gangaagene tntgaaggee
                                                                           480
aaaacagagt acanaagttt conngttggc ttggattttg aaaattcnng aattntntat
                                                                           540
aacgggcttn tttaaaaagg atnggnttan gnacctttnt taaat
                                                                           585
      <210> 77
      <211> 591
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (591)
      <223> n = A, T, C \text{ or } G
      <400> 77
ggtacgcggg agtcatattt atgaaaaaag gtttgtgttt tactcttgct agtgagaaag
                                                                            60
tgggacaaaa tatacttttg aaataaaatg ctatatggca cctaattatt ttttctttta
                                                                           120
aaatgeetta agttgeagte teattttgat aateatttge tteeagtgtt taaaaattaa
                                                                           180
aaaaagaatg gggagaaggt tatgagaaga gcattattaa gtttccaaat ttaatttgaa
                                                                           240
ttccaaattc acctagcaat aaaatctaat ttttaaaaaag tatataaata taaaatgtat
                                                                           300
aaatgatgga tagatttttg tattgatttg caaaatgcag attatatttg ataggctata
                                                                           360
gtatgtagat attectttta ggaatattac agetgtaaat tatatgagae ttgccagtca
                                                                           420
aatgctattt ggtttaaaaa aattattgca atctcaagtt aatggaatat ttttaaatcc
                                                                           480
cacattcaga gitaaaacct ngttttcaat gggtttttan tgtggcactt gnttatagat
                                                                           540
taatttttaa taacctgttn ggaancnggg cettttaact ggteettggg g
                                                                           591
      <210> 78
      <211> 252
      <212> DNA
      <213> Homo sapiens
      <400> 78
actgagaagt attttcagtg attcgaccca gaccagattt caacacatgg ttcccataca
                                                                            60
ggaaggactg ctctgcacca ggctttatcc aaactttata cttggcataa ggtgcaaggt
                                                                           120
aatccagage tgtgacgtge aaccgaaact tgtgggtttt agtgaatttt ccaaagcagg
                                                                           180
tecceagega caccagettg tecceggaaa tattggegge cagetteata atetteteae
                                                                           240
tcacatagta cc
                                                                           252
      <210> 79
      <211> 571
      <212> DNA
      <213> Homo sapiens
```

<220>

```
<221> misc_feature
      <222> (1)...(571)
      <223> n = A,T,C or G
      <400> 79
gctcgggcaa gcactttaac cttttaagcc caaccagatg agttgcctgc agttttggag
                                                                         60
geetteagag cattteacta gacetetgte tgtgteggte eagtgtettt ageeaagett
                                                                        120
tgattaaaga tgacttcctt gtttgctcaa gaaattcgcc tttctaaaag acatgaagaa
                                                                        180
atagtatcac aaagattaat gttacttcaa caaatggaga ataaattggg tgatcaacac
                                                                        240
acagaaaagg catctcaact ccaaactgtt gagactgctt ttaaaaggaa ccttagtctt
                                                                        300
ttaaaggata tagaagcagc agaaaagtca ctacagacca ggattcaccc acttccacgg
                                                                        360
cctgaggtgg tttctcttga actcgttact gggcatcagt agaagaatat attcccaaat
                                                                        420
ngggacaagt tettttagga agaceetta teettttget ggtgaaaate aaaatgaage
                                                                        480
nnaaaatccc ttcaaaatga ggccaacgan taacttttt aaatggcttt tcaaaaagcc
                                                                        540
ntgttaatta ancttnantg taaaggnttt t
                                                                        571
      <210> 80
      <211> 595
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (595)
      <223> n = A, T, C or G
      <400> 80
acctetteet gttegaatgg gttatecagt aaaaaaggge gtgeecatgg caaaggaggg
                                                                         60
aaatctagaa cttttaaaga ttcccaattt tctgcatttg actcctgtag caattaaaaa
                                                                        120
gcactgtgaa gcccttaaag atttttgcac tgagtggcca gccgcactgg acagtgacga
                                                                        180
gaaatgtgag aagcattttc caattgaaat tgacagcact gattatgttt catcaggacc
                                                                        240
atctgttcgg aaccccagag cacgagtagt agtctcaaga gtaaagcttt ccagtttgaa
                                                                        300
tttagatgat cacgcaaaga agaaattaat taaacttgta ggagagcgat actgcaagac
                                                                        360
cacagatgtg cttaccatca aaacagatag gtgcccttta aggaggcaga attaccatta
                                                                        420
tgccagtgta tctactaaca gtgttatatc atgagtcttg gaatactgaa gaatgggaaa
                                                                        480
aaagttagac tgaagccgac ttggagaatn tatatgggaa aatactatca gaaagaaata
                                                                        540
tctggnaacc cttttccgat gaaagtgctg anaaaatntg gaattaataa gaagn
                                                                        595
      <210> 81
      <211> 601
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(601)
      \langle 223 \rangle n = A,T,C or G
      <400> 81
acgcggggga aaacaagatg gaggattcgg cctcggcctc gctgtcttct gcagccgcta
                                                                         60
ctggaacete cacetegaet ceageggeee egacageaeg gaageagetg gataaagaae
                                                                        120
```

```
aggttagaaa ggcagtggac gctctcttga cgcattgcaa gtccaggaaa aacaattatg
                                                                         180
ggttgctttt gaatgagaat gaaagtttat ttttaatggt ggtattatgg aaaattccaa
                                                                         240
                                                                         300
qtaaaqaact qaqqqtcaqa ttqaccttqc ctcatagtat tcgatcagat tcagaagata
tctgtttatt tacgaaggat gaacccaatt caactcctga aaagacagaa cagttttata
                                                                         360
gaaagetttt aaacaageat ggaattaaaa eegtttetea gattatetne etteaaaete
                                                                         420
taaagaanga atataaatco tatgaagooc aacttoogno ttotgagoag ttttgaatto
                                                                         480
ttncttactg atgccagaat tangengnte ttaccettac tcattgggag acatttctat
                                                                         540
caaaqaaaqa aaqttcaqta tctqtaaacc ntttgtccaa gaatttttca ggagagatca
                                                                         600
                                                                         601
      <210> 82
      <211> 606
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(606)
      <223> n = A,T,C or G
      <400> 82
cgaggtactt tgaatatgga gtagtttaca gctatttttt tttcttactg gtaatcttaa
                                                                          60
ctaatatgat teeettatta gagageetet caeteeeca eeeecaaaaa tgtetaetat
                                                                          120
tcatgacagt aaccaattat tctqqacaaa ttgcttcttt ttaatttgag ctatctgcca
                                                                         180
tggactttct aaaatggaaa cacagcctga gtgtatctta gggagagttt gattgaaaaa
                                                                         240
atccaaatca ctatccatat agatcatgga tataaagaga tacctgattt ttattaaaaa
                                                                         300
gatacttttt caaatttaag agttaatctt ggaaatttgg aacaagtaaa ggggcaagta aaccttttga tgaaatataa aaggactcat tgcatgaagt gactatcaaa ttctgngatg
                                                                          360
                                                                          420
tgnggcttet taaaaatatt etcagggett tgggggeetg ccanatggta eetgeeegge
                                                                          480
ggccgtcaaa agggcgaatt ccncacactg ggggccgtac tagggggtcc caacttggac
                                                                          540
ccaacctggn gnaaataang gcataantgg teenggggga aatggtnnee gttecattne
                                                                          600
cccann
                                                                          606
      <210> 83
      <211> 613
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (613)
      <223> n = A,T,C or G
       <400> 83
gegtggtegg geegaggtac acgttegtea tggeggetgg ceetggacet gggtaggggg
                                                                           60
teegggttea gtggtaatag eggeggagat gggggageet eegettgget tettteacae
                                                                          120
gggttgcttc ggaggaatcc qccqtqcaaa tctqtccqcc cccttggcca ctgatccccc
                                                                          180
gaagagette tgtegeeget etaggaatae agacattgaa gtttgggaea agatatttat
                                                                          240
ctaacttctg tgtcaaaatt agcgacctgc tatggcaatg aagaaagaaa ctgaatttgt
                                                                          300
cattttcacc tgaagaaaaa tgatagacaa aaatcaaacc tgtggtgtag gacaggattc
                                                                          360
tgtgccctat atgatttgct gattcacata ctcgaagaat ggtttggtgt ggaacanttg
                                                                          420
gaggactatt tgaattttgc aaactatctc ttgngggttt tacaccacta atacttttaa
                                                                          480
tacttectta etttactate tttettetet acettactaa taatttetta eacattatta
                                                                          540
```

```
agaagaaaga tgttttgaaa gaagcctact ntcataatta tnggatggtn caagggaaac
                                                                              600
anggcactnt ntg
                                                                              613
       <210> 84
       <211> 605
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (605)
       <223> n = A, T, C \text{ or } G
       <400> 84
ggtactatct gctgctggca aatggggttg ctctgggtga cagggatctg ctgacccaat
                                                                               60
gctatggttt gttccagtca atgagttgag aaggctaaag ccttggttcc tatcattctt
                                                                              120
catcactaca ttggaccaca cattggcatt cagggcttgg acaattcgct ttactcctgt
                                                                              180
agattetggg aagteateat ecteeteagg caacteetet ggaetaagtt etaceaatte aaageeatgt ttgaggeace attettgage tttttgtegg tttataceat etteagaeae
                                                                              240
                                                                              300
totategeag accaagatea teaceteagg taaccatget tttgecagtg gaagecatga
                                                                              360
ggagacacta tcaaggcccg atttttgtgt gctgtcaaag taaaccacaa atgcttggac
                                                                              420
agattetgea atetetgeag taaccagaaa tttgttggge accecacata gattgagtet
                                                                              480
gctgaaaagt atttattatc aatggncccn ggataaaact acacattatt tggaagtact
                                                                              540
ttencaataa gaaettntgg teeaaggtat ttttggaeen aanggnetet tgaaaaaaeg
                                                                              600
gagga
                                                                              605
      <210> 85
      <211> 603
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (603)
      <223> n = A,T,C or G
      <400> 85
acagggaatg aagactcgaa gaggagatgt cactttcctg gaagatgttt taaatgagat
                                                                               60
tcaattaagg atgctacaga acatggcttc aattaagaca actaaagaac tcaagaaccc
                                                                              120
acaagagact gcagagaggg tcgggctcgc agcactcatt attcaggact tcaaaggttt
                                                                              180
actettatet gactacaagt teagetggga tegtgtttte cagagtegeg gggacacagg
                                                                              240
tagagtaaac tgcanagetg cetgtetgtg acttecaagg etaggteata aaaggagata
                                                                              300
aagettette tggetgggtg ggetgettge tettgaacet teagtetatg cacgeaacat
                                                                              360
gcctttccag ccttctgtgg ttgtagagtg natagaaagc aattggatca ctatngacag
                                                                              420
cggggtaaaa cttgaggaag caacctccgc caggnggtac atggagganc cctgaannaa
                                                                              480
aggaanaaaa gggcacangg gcttaatcct gtcttggaat gcttncctnt gcaatggnnc atttcaatgg ccnagccaat tatgccatcc ctgcnttaan accatgggcc ttcnttgnca
                                                                              540
                                                                              600
ttn
                                                                              603
      <210> 86
      <211> 583
      <212> DNA
      <213> Homo sapiens
```

```
<220>
      <221> misc_feature
      <222> (1) ... (583)
      <223> n = A,T,C or G
      <400> 86
actgtaggta tttattaata atagcaatga agatgaaaga gtgatgtatc agagaggtgg
                                                                      60
agataaaatc agtaaaactt agacactaaa tgatagggga aggtggagga gaggaatgaq
                                                                     120
cctagaaaac ttagaatata atggttctaa aattaaccaa agtaagggac acaqqcatta
                                                                     180
gagtaggttt tgcagagaat gaatgtttta agacacacac aggtgtctct gggacaacca
                                                                     240
agaaaagtgc aacaggcaga tggattgagg agtctggcta aagataagga tttaggaact
                                                                     300
360
                                                                     420
caggagtgac ccaaatgtag aagtcaggga ataaacttta aagtnggggt gtcaaaatgc
                                                                     480
naatccgaaa aaaagtnagt nccttggccg gaccccctag gcgaatccac ccctqqqcc
                                                                     540
gtctanggat ccacttgncc aacttgggaa nntggctnct ttt
                                                                     583
      <210> 87
      <211> 332
      <212> DNA
      <213> Homo sapiens
      <400> 87
acgcgggggc attgctagaa gccggcagga gtgactctcg gcatggagga cccatctcct
                                                                      60
agcacacgtg cccactgaag tggcaccaac agaagtttgg cttgaactaa aggacatttt
                                                                     120
attttttta ctttagcaca taatttgtat atttgaaaat aatatatt attttaccta
                                                                     180
ttagattctg atttgatata caaaggacta agatattttc ttcttgaaga gacttttcga
                                                                     240
ttagtcctca tatatttatc tactaaaata gagtgtttac catgaacagt gtgttgcttc
                                                                     300
agactattac aaagacaact ggggcaggta cc
                                                                     332
      <210> 88
      <211> 592
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(592)
      <223> n = A, T, C \text{ or } G
      <400> 88
cgaggtacgc ggggacaacc agctgactcc cgtagaggaa gacactgtgg aggccagttc
                                                                      60
tggagctatt gcagcctcgg ttgcccggcc cgggacccga acccgaaaaa gttatcgtca
                                                                     120
gaatgteggg caaagaeega attgaaatet tteeetegeg aatggeaeag accateatga
                                                                     180
aggetegitt aaagggagea cagacaggte gaaaceteet gaagaaaaaa tetgatgeet
                                                                     240
taactetteg atttegacag ateetaaaga agatnataga gaetaaaatg ttgatgggee
                                                                     300
aagtgatgag agaagctgcc ttttcactag ctgaagccaa gttcacagca ggtgacttca
                                                                     360
gcactacagg tattccaaat gtcaataaag ccccagtgaa gattcnagen aagaaagata
                                                                     420
tgtacnagtg gtactttgnc ngtatttgaa cattccntga aggactgcng gttttnactg
                                                                     480
cttgggttaa cccaagtggg gacnnnttgc ttaaattaaa gaggaatttt gcccaancnt
                                                                     540
gggacttetg gnggaattac ttttttggaa actttttggn accttggagn aa
                                                                     592
```

```
<210> 89
      <211> 630
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(630)
      <223> n = A, T, C or G
      <400> 89
acgcgggggt ctttgggccg gcgcgaacca tggccggcat ggtggacttc caggatgagg
                                                                         60
agcaggtcaa gtcctttttg gagaacatgg aggtggagtg caactaccac tgctaccacg
                                                                        120
agaaggaccc ggacggttgc tatcggctgg tggactattt ggaagggatc cggaagaatt
                                                                        180
ttgatgaggc tgccaaggtg ttgaagttta actgtgaaga gaaccagcac agtgatagct
                                                                        240
gctacaaact gggggcctac tatgtgactg gaaaaggtgg tctgacccaa gacctgaaag
                                                                        300
ctgccccagg tgctttttga tggcgtgtga gaaacctgga aagaaatcaa tagcancatg
                                                                        360
tcacaacgtt ggccttctgg cacatgatgg acagggtaat gaagatggcn acctgacttt ggaaaaaggca aggactacta ccaaaggcct gngatggngg ntatctttca gtgcttnaaa
                                                                        420
                                                                        480
cctaatgcat ttttcttcag ggggcccaag ctttccaagg acatggcctt gcctgtnaat
                                                                       540
cttcattaaa gccttgacct ggtcatattt ggccttgcca tgcaatccat ttacttggcc
                                                                        600
ggacacctan gggaatcacc actggggcgt
                                                                       630
      <210> 90
      <211> 653
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(653)
      <223> n = A,T,C or G
      <400> 90
60
ataaaataaa gcaaaaatat aaaatgttaa aaaaaaaaca aaaaaaggga aaaaggaagc
                                                                       120
tgattgcctt ggtgagtcaa cactgggtat tttctgacca ctatttgaaa caaaaaagga
                                                                       180
aaccactgat attetatgca aagatetgtt cetggaagge actetgegga gacaccagga
                                                                       240
gaacttttat caatcettca ttgatttgaa gtaaaagtge taaagcaatg gttggtgggt
                                                                       300
ggcaacccat tagcagatca caaaatcact gtagtgggta actaaacaag aggaaacaca
                                                                       360
agacggcate etgtgtaact ggggttaage attactetet gaaactcatg geatcagttt
                                                                       420
cetettagge tetteccaca aagtataate atgtteattt cagtttacaa teeettgeag
                                                                       480
tcccatcgat ttgtgagaat atcccaagtc atncacagng gagnetggaa atggtentan
                                                                       540
ttgtcctgcc cggcngccgt tcnaanggcg aattcaacac actggcngcc gttctaatgg
                                                                       600
atccaactcg naccaacctg gnggaacatg gctactggtt ctggngnaaa tgn
                                                                       653
      <210> 91
      <211> 657
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
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<222> (1) ... (657)
      <223> n = A, T, C \text{ or } G
      <400> 91
actititit tittititt tittititig ggagaaaagc cinactceqt tqcccacqtt
                                                                            60
ggagtgcagt ggcgtggnca tagcttattg catgcagcct naacctccca ggctnaaqca
                                                                           120
atnotecnae etnnnegtge tennttnntg gaactaenea theachenat tatgeceane
                                                                           180
tnqtnqttgt naatttaaag tganaccatg cncncaggnn gnatggcntt nnntancnan
                                                                           240
catgcatgct cagetgtgta gtgcacgcac aggataaatg gaagggggat ttgatcaggg
                                                                           300
tttttgtcac atnagcattn naaatccgna ngactgccnt gtgtctgcct ttgnaagggc
                                                                           360
ctgggagtat tctgtgtagc ctttgnaaat aagggnaaaa tgngcncctg ccaaagaagt
                                                                           420
entigetact nigggigngt caaaatnice eigtaactig teaatggnea caagetiggn
                                                                           480
ggngtntttg ggntcttggn tgtcnttttn acgtctattg nccatgtggt tcctatatga
                                                                           540
cacantecte ntnataatee ntganaattg ctaanntgen etttttttt ttttnanatt
                                                                           600
nattttgctn ttaaantagc ttaanncttt ntttatcctn gggcanccna anncaat
                                                                           657
      <210> 92
      <211> 653
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(653)
      <223> n = A, T, C \text{ or } G
      <400> 92
accataaaac cattaaaagc aataaataac tagagtcatg tgagatgttt caaagactgc
                                                                            60
tggaggtttc tgtaaaccag ggtaatcaga aatattaccc ttgtagatag ccctctcata
                                                                           120
ccagtaaata caaagagtta aaattccaat gccacagtgt aacagttaac aatctatttt
                                                                           180
gtaattttaa atattactac attaattcac cctgagaata cagaggaaac atttaataca
                                                                           240
agacattctg atatgntttt ttttcccatt gnatttgctt tcttctggnt ttcatcagcc
                                                                           300
ctttaagggc acagatattt taatttaaag ggtgatttgg atatgctttt ttggtaactg agatttatgc cacagtcaga tactggtgat agaaaagccc aaaaaggntt gnagaaaaga
                                                                           360
                                                                           420
ggcaagcagc aatccccagg cagaaaagac ngaaagtctt gaaaaagaag aggagtaaaa
                                                                           480
atttttttaa getgnteaat geeetgtatt tgggnacaag tacetttatt tteetttage
                                                                           540
tganggnant cagagtaacc gaattggnag nnnactattt tcnctggnaa ggaaaataga
                                                                           600
atttggnaat cccnggaang gtnctngaaa tnnagcccca tccatttgnn gng
                                                                           653
      <210> 93
      <211> 640
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (640)
      \langle 223 \rangle n = A,T,C or G
      <400> 93
acagagaaac cacaggttgc cctttccaca gctggataga cttatccaaa acggcaggat
                                                                            60
ggttctgtat taatcttttt ggaaagcatg tctgtattaa gattgcaaaa catacagata
                                                                           120
gctaccacaa attaggtcaa acgactgatc aagttgtaac atctgtgagg tcaaattcca
```

180

recommendation of the same

```
aagtgcctag atacacattt atacaacaga ccataagagc tgaattcttt acaaatgtct
                                                                           240
ttatgggcat gtaaaattga ctctgcattt ctgcatgtgt gcattccata agagagacca
                                                                           300
gtotgcactg agtoatatat actocaactt gaaaaagtaa gtgnaacaac tggntaatca
                                                                           360
tgcaagtctg gttgnaatat aacaatgact ggnaaaacat gaattcttcg cacagtagta
                                                                           420
ataggngcae tnatttaaaa ccctnccgaa aaacctgnat ttggtgcaan atctganttt
                                                                           480
aagnggtagt aacttgacnt ttaaaaatag tttgaacnat ttanaaaggn aagccaactt
                                                                           540
ttacttaaaa gaatcccaag tggnaaaanc tggntttcaa tggaatgaac tngqtqnqac
                                                                           600
ctnccctaat nngaccttga gcctatnagc taatntangg
                                                                           640
      <210> 94
      <211> 658
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(658)
      <223> n = A, T, C \text{ or } G
      <400> 94
acgogggcca agottttttt ttaatttggt gttttctccc atcotttccc tttaaccctc
                                                                           60
agtatcaagc acaaaaattg atggactgat aaaagaacta tottagaact cagaagaaga
                                                                          120
aagaatcaaa ttcataggat aagtcaatac cttaatggtg gtagagcctt tacctgtagc
                                                                          180
ttgaaagggg aaagattgga ggtaagagag aaaatgaaag aacacctctg ggtccttctg
                                                                          240
tecagetete aageactage ettacteage tatecattat agetetegee ttaagaaage
                                                                          300
catgattaac ttatgaaaaa attatttggg gacaggaatg tgatacette cttggntttt ttttgcaane ctcaaateet aactteetge eccacaatgg tgagcaggtt eccetgatae
                                                                          360
                                                                          420
ttetttett taatgattta actatnaact tgnataaata acttataggg gatagggaaa
                                                                          480
attectgaat tecagaatge catetgntaa aaaagaatnn aaatgggaag tnggaetnaa
                                                                          540
aaggagccaa cagcatgctg cggtggnngn cacttctttg cnctatccca ggaaggaagg
                                                                          600
tececatttg gaaaggggtt ettneteaet ggnaceggtt tgaentnatt ggnaenee
                                                                          658
      <210> 95
      <211> 392
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(392)
      \langle 223 \rangle n = A,T,C or G
      <400> 95
actcagactt gatcgattaa tgaagtggtt attttggcct ttgcttgata ttatcaactc
                                                                           60
actggtaaca acagtattca tgctcatcgt atctgtgttg gcactgatac cagaaaccac
                                                                          120
aacattgaca gttggtggag gggtgtttgc acttgtgaca gcagtatgct gtcttgccga
                                                                          180
cggggccctt atttaccgga agcttctgtt caatcccagc ggtccttacc agaaaaagcc
                                                                          240
tgtgcatgaa aaaaaaagaa gttttgtaat tttatattac ttnttaagtt tgatactaag
                                                                          300
tattaaacat atttctgnat tcttccaaaa aaaanaaant aatnaantta naanctttta
                                                                          360
aanatanaaa taaaataata angaccattg ag
                                                                          392
      <210> 96
```

<211> 655

```
<212> DNA
       <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(655)
      <223> n = A, T, C or G
      <400> 96
ggtacaggtt tttatgtgaa catacatttt cattttctgg gataaatgct caaaagggca
                                                                           60
actgttgggt tgtatggtaa acacatatat ttttgtaaga aactacccta ctcttttcc
                                                                          120
agagtggctc tactttttac atacagccac tcatacaatt cagacagcaa tgtatgattg
                                                                          180
atccagttte tteacateet caccageatt tggtattaet actatttttt atettaacea
                                                                          240
ttcacataga tgtgtgtaat gataccacat gtggttttaa tttgcatttc caatggctaa tgatgttgag tatctttttg tgtgctaatt tgccatctat gtatcctctt cggtgaaatg
                                                                          300
                                                                          360
tetteatgte tittgnetat titetatita agneatitgg tettittaet attgagtitg
                                                                          420
agagggtttt tatatateet agataaaaat eetetggtan anatgtgggt geetggaatt
                                                                          480
ttaacataac ttctacccan ggaaaataag taaaatttcc accettgetg genageetta
                                                                          540
cttaatnccg gccttaangg tccttctaga gaattaagaa gatttgaggt ttaaatanaa
                                                                          600
tcagggentt aaaaagtaat cctaaaaten ggtttaagea agecatatee tgggg
                                                                          655
      <210> 97
      <211> 224
      <212> DNA
      <213> Homo sapiens
      <400> 97
acaagtttaa ggtaggacgc agcattttat agtgttacgt ccttcctcc cacattcctg
                                                                           60
tgaggcggaa caagaacaat tacttgaccc tggaggaaga cgacgccttg tggtcaggga
                                                                          120
gagaacagca gttcatgctg gctgcctcgt ctttccaggc ctgctgctgc ccaggcttct
                                                                          180
actgaccttg ttaggtctga ttctagaaaa tgaaggcagg tacc
                                                                          224
      <210> 98
      <211> 582
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(582)
<223> n = A,T,C or G
      <400> 98
ggtaccacca tgcctggttt attgttttat ttttttggca gagatgggtc tcactgtgtt
                                                                           60
geccaggetg ateteaaact cetggeetea agegateete ceateteage eteccaaagt
                                                                          120
getgggatta cagacetgag ecaccacace tgggcaacag agtgaaacet gteeetgttt
                                                                          180
teetgetett acteteacet etgaggeete etetgeetgg aagagattae agggaaatte
                                                                          240
caggcagccc ttgtcaattg tttttatgaa ttctttacct gttcctttta aagacaagga
                                                                          300
aactgaggcc caaagttcta agttgtttgg caaatggagt ctcctaccct cagctcctgc
                                                                          360
aaggacctgg gggaccccca ggtccagcag ccacatgatt ctgcacagac agggacctag
                                                                          420
agcacatetg gatttaagce caccetggca actggetget agagactnee aagatgeega
                                                                          480
taataggatc tgccnttaaa aaatctggat tctggcctgc ntaantgcta cttcatttgg
                                                                          540
ctacaaagnt ttaaggngga accnttaaaa ccttccccaa aa
                                                                          582
```

```
<210> 99
      <211> 619
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (619)
      <223> n = A, T, C \text{ or } G
     <400> 99
ggtacagtgg tccttttcag agttggactt ctagactcac ctgttctcac tccctgtttt
                                                                         60
aattcaaccc agccatgcaa tgccaaataa tagaattgct ccctaccagc tgaacaggga
                                                                        120
ggagtetgtg cagtttetga caettgttgt tgaacatgge taaatacaat gggtateget
                                                                        180
gagactaagt tgtagaaatt aacaaatgtg ctgcttggtt aaaatggcta cactcatctg
                                                                        240
actcattctt tattctattt tagttggttt gtatcttgcc taaggtgcgt agtccaactc
                                                                        300
ttggtattac cetectaata gteatactag tagteatact ceetggtgta gtgtattete
                                                                        360
taaaagettt aaatgtetge atgeageeag ceateaaata gtgaatggte tetetttgge
                                                                        420
tggaattaca aaactcagag aaaatgtgcc catcangaga acatcataac ccatggaagg
                                                                        480
atnaaagccc caaatggngg naactgataa tagccctaat ggctttaaga atttgggcac
                                                                        540
actnttacct aggngaaccc atttgancen anggggetta aaggettntt acttcaactg
                                                                        600
aaagttnagg qaaaaaaan
                                                                        619
      <210> 100
      <211> 614
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (614)
      <223> n = A, T, C or G
      <400> 100
acgcggggga agcaaaggag agggaagctg gaagcacctt tggcccggga cagaaatctg
                                                                         60
gagagettgg ctacetecat ceteeteagg eeggageagg etteetgaga gagteeaggt
                                                                        120
cgtaggagtt ttacgactta gaaaagcggg ctgcagattc cttcctgggt gtttggttca
                                                                        180
agccctggct ccagcctcac tctcagtctt cccgggagtt cgtgggattt ggaccttaga
                                                                        240
ttattagtat tattttgagg gcctcctgtg tgtaagcact ggttgtgcgc agatggctgt
                                                                        300
gcagagggcc atgaggtaga ggctggggaa atgagggctt ggaggtgctt gaggtatqgt
                                                                        360
ctttacctac gtgaaatgtt ggaggttgag atgaaaactc ttgctttgaa atcttcatgg
                                                                        420
aggactacat catttcaatc ctgaatctgg ctcaattcta ttaatcactt aatacctgga
                                                                        480
ttaaaaaacg nttaantggg ccaggencaa tgggtcacgc ctgnaatccc agcentttgg
                                                                        540
gaggccaagg cangccggat acnttagggc ngnanttnaa accancttgg caaattggga
                                                                        600
aacccgcntt tntn
                                                                        614
      <210> 101
      <211> 625
      <212> DNA
      <213> Homo sapiens
      <220>
```

```
<221> misc feature
      <222> (1) ... (625)
      \langle 223 \rangle n = A,T,C or G
      <400> 101
ggtactttgc ctacggcagc aacctgctga cagagaggat ccacctccga aacccctcgg
                                                                       60
cggcgttctt ctgtgtggcc cgcctgcagg caagaagggg ttaaaagtgg aatgtatgtt
                                                                      120
gtaatagaag ttaaagttgc gactcaagaa ggaaaagaaa taacctgtcg aagttatctq
                                                                      180
atgacaaatt acgaaagtgc tcccnnatcc ccacagtata aaaagattat ttgcatgggt
                                                                      240
300
                                                                      360
                                                                      420
tatttttaac acttgagaac cagggatttt gggggattct ccaacgtttg ttcaatttta
                                                                      480
agaantggtt tgaaggagtt ttttacttgg gtnattcntg gttttaggat tttnnanngn
                                                                      540
aanntggntt nggngtttgn nnttttaann gggntntttt ngggtcttna aatttttcca
                                                                      600
anaaanngtn gnttccttcc cqqnn
                                                                      625
      <210> 102
      <211> 605
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (605)
      <223> n = A, T, C \text{ or } G
      <400> 102
ggtacaagaa agaaaaata taaaaacaag tctgctgagt gtcgggagtt ggtgagggat
                                                                      60
atcctaccat attgtgacgg agtccaaata gaaaacatgc agcaacagtt ctcctgcttt
                                                                      120
atcagctccc tggaaaataa accagtaacc ctggtagtgc agtaaccatt tggttaacag
                                                                      180
gacaaacttc ctgatggaca cagatagtaa ttcactgcat ttcccttctc taacttctct
                                                                      240
cttcacacca attccttttc tttcctttaa gatgggtttc atcctgttga caaaagattt
                                                                      300
ggttttattt gtaaagtaaa gcagataata tcctgattga agtattcaat gatttaattg
                                                                      360
aggatgettg gggatcaaac tttgtaaaaa ggtcaattaa gctagttagc agagactatc
                                                                      420
agtggcttgc agaaaaaaa ntcngatata tggtttggta aaangcccaa aggataaccg
                                                                      480
ngaaaaatcc tanggatacc gggacctaat taatcaaagc canaggggga ccttggttaa
                                                                      540
ancenttact tnggggangg getnaanggn ggntecaaac naaattggtt cecaacggge
                                                                      600
ccggg
                                                                      605
      <210> 103
      <211> 251
      <212> DNA
      <213> Homo sapiens
      <400> 103
acgcgggatt ttacattcca tcttttctga agattgtcct acaatttgga ttttgatcat
                                                                      60
gacaaagaag attaaaattt cattagcatg aatgcaattt gttaaagcag actgatttgt
                                                                      120
ttctaagata tttttggttt ttttaaaact gataataatg ctgaattatc ttaagtgaga
                                                                      180
tgttaagece actttgttet tttaatgtaa tggagettat gggtagaaga ceatgtetae
                                                                      240
taattacaaa a
                                                                      251
```

```
<211> 293
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (293)
      \langle 223 \rangle n = A,T,C or G
      <400> 104
ttaatcttgc acaaatggca ttttattaaa gaaaatctaa tttacaaagc tttgtaaatt
                                                                          60
ttaagaaaaa cattcataga tcataaacaa aaatttcaat atgcaatatt caaatttaca
                                                                         120
agaaaataag cacaaacttt tagacagtgc agttattgct gcactccttt aattccttat
                                                                         180
ccagagccca aaaaatgtag acaaacccta aaaatgtagc agaagcattt ccgcacactg
                                                                         240
gtgtccagaa tctagtttgt gcanaaatgt ttccactaga tttatagagt acc
                                                                         293
      <210> 105
      <211> 586
      <212> DNA
      <213> Homo sapiens
      <221> misc_feature
      <222> (1) ... (586)
      <223> n = A,T,C or G
      <400> 105
acatcatttc tgccatgtgg gacattttct tgggaatata caagtaatac tccatgtagc
                                                                          60
ctgacaggtc ctcaatggtc acatcatcca cgaagactcg agcttgctca gaacaggatc
                                                                         120
ggggagagcc agacagagtt ctggcgtgca gcgactgaga gtagtcctca agtgtggatc
                                                                         180
ttegttetgg agecaaggga gggacaetet gegggeetga aaaggaatae aetteeatat
                                                                         240
catgccatct cttacactgg cattccttgc ctatgcatgt gcatggcttg ccctggttta
                                                                         300
gettggaaac tgattgaaag teagagagat eactggettt gagaettget tgggggaett
                                                                         360
gggtagccgt cagaggagtc ttccttctta ctctctgatg ggagccttgg aacagaaagt
                                                                         420
tetcaaange tnaacgaetg geeetggggt gaatageate gagagaagta nacettette
                                                                         480
ctgnactgaa ctnttaaggg gatgaaattc ccagccaatg gtggccttan gnnangcaan
                                                                         540
ntggcttttg gcttgaatta ctggntggaa aaaacctttg gccntt
                                                                         586
      <210> 106
      <211> 644
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(644)
      \langle 223 \rangle n = A,T,C or G
      <400> 106
ggtacnttga ttgctcanat ataangaaat ggcccaatga acgtggntgn gggaggggaa
                                                                          60
anangaaaca gagctagnca tatgtgaatt gntctgtgnn ataaacatgt taaaacanac
                                                                         120
aaanatggnt attittettt neeteeggae agtgeacatt ateatnigaa etaeetgggg
                                                                         180
attctntatc anaactggtc ttgttgaata tttatactta attgaaataa ttccttanng
                                                                        240
```

```
gaggentgtt taaaacgtat taacaggana ttgtgtntna nacatttaat gaaanacgaa
                                                                               300
attccacnag aatganntaa gtcactttcc aagtgggtgt cattttgtta aaccctngtt
                                                                               360
tacctgtttt gctattntta contttcatt tggaangatg ntttgagntc gtanttacca
                                                                               420
gggnaaagac gggttncttc ctngctgnnn cttnagccnn tgctaaaaag cnttaatttt
                                                                               480
ntgcnattng gnncttcctg ctggtaatcn tggaaaaant gggnnaantc cagctttntt tnttggcngc ccaaaaangg attcnnantn gnnannnaac ctttggttcc ntaannaana
                                                                               540
                                                                               600
aaangtatnc anaangaacc ttgncatgcc ngccnntnta aang
                                                                               644
       <210> 107
       <211> 618
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(618)
       <223> n = A, T, C or G
       <400> 107
ggtacagact tgccctttga aatctatacc tctggataca ttagaggcat tttattaaca
                                                                                60
aaggcccttc taaatgtgct atttatttga caataactat cagatttgcc ttaattttqt
                                                                               120
gtttatagca tttatcaaaa cgtatcctca tagactttat gcagattaat atggtcaatt
                                                                               180
gatttggata aaagaaagta atttcagggt ttgtttttaa gccaggacaa gaagtgcaaa
                                                                               240
tgcctctttg aagcaattta ggctaaactg attttgaaat ttcaaaatgt tttattttac tttgttttat taagccagga caagaagtgc aaatgccctc ttttgaagca attcaggcta
                                                                               300
                                                                               360
ggtaaacccg attttggcca tttcaaaacc gtttaattta ctttggttta atatcaqaqt
                                                                               420
cttataaaac tgntgncaaa aatttctgaa ggctttngaa aagggttggt agtggaccct
                                                                               480
gcccgggcgg ccgntcnaag gcgaattcag ccactggcgg ncgtactagg gatnccactc
                                                                               540
ggacccanct tggcggaatc atgggcataa ctggttcctg ngtgaaatgg gatccgttac
                                                                               600
aattcccaca acatanng
                                                                               618
       <210> 108
       <211> 620
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(620)
       <223> n = A, T, C or G
ggtaccaaag gagaatttgg agagctggct aaattatttg aagaaagaat tgccaacagt
                                                                                60
ggtgttcaga gcctcaacaa aaccaaagga taaagggaag ataaccaagc gtgtgaaggc
                                                                               120
aaagaagaat gctgctccat tcagaagtga agtctgcttt gggaaagagg gcctttggaa acttcttgga ggttttcagg aaacttgcag caaagccatt cgggttggag taattggttt
                                                                               180
                                                                               240
cccaaatgtg gggaaaagca gcattatcaa tagcttaaaa caagaacaga tgtgtaatgt
                                                                               300
tggtgtatcc atggggctta caaggagcat gcaagttgtc ccctttggac aaacagatca
                                                                               360
caatcataga tagccccgac cttcatcgaa tctncactta attccttctt tgngccttgn
                                                                               420
ttttgcnaag ttcanccaag gttttgaagt antaaaance gatggaaget tgccantgce
                                                                               480
atcettteca agettgatge ttgacaggta gtanettgne egggeeggee gttenaaagg
                                                                               540
gcgaattcaa cacactggen gccgtactat ggatccgage ttggnccaaa cttgcgtaat
                                                                               600
catggcatnc tggttcctgg
                                                                               620
```

```
<210> 109
       <211> 317
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (317)
       <223> n = A, T, C or G
       <400> 109
tttgtatttt tagtagaggc agtgtttcac cgtgttagcc aggatggtct cgatctcctg
                                                                             60
acctegtgat ccacccacct cgacctecca aagtgetggg attacaggeg tgagecacca
                                                                            120
cgcccggcct ctttttttt tagctgccaa tctttttgaa ggaatattct tacctctact
                                                                            180
ttgtcacctt ctactggctc cttaactaaa atctgccatt tggctctctg gttaacagtc
                                                                            240
cetteetgta aagtetaaaa tettaattet aaateeacag titaatteac aagetagtae
                                                                            300
cttggccgng accacgc
                                                                            317
      <210> 110
      <211> 603
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (603)
      <223> n = A,T,C or G
      <400> 110
ggtacattca ggatccctcg gccaaggact ggaccagaag aacacttggg aatcttgggt
                                                                            60
ccacttatca aaggtgaagt tggtgatatc ctgactgtgg tattcaagaa taatgccagc
                                                                           120
cgcccctact ctgtgcatgc tcatggagtg ctagaatcta ctactgtctg gccactggct gctgagcctg gtgaggtggt cacttatcag tggaacatcc cagagaggtc tggccctggg
                                                                           180
                                                                           240
cccaatgact ctgcttgtgt ttcctggatc tattattctg cagtggatcc catcaaggac
                                                                           300
atgtatagtg gcctggtggg gcccttggct atctgccaaa agggcatcct ggaaccccat
                                                                           360
ggaagaccga gtgacctgga tenggaattt geattggtgg teetgaattt tgatgaaaat
                                                                           420
aancetggna tttggaagga aatgtgcaac catgggteca agaatecage ennattaace
                                                                           480
taccggatga acctttnttg gaaaccataa aatgcctgca atcaatggga actttttcca
                                                                           540
accttanggg cttaccatga ccttgcccgg ccggccnttt aaanggccaa ttccacccc
                                                                           600
tgg
                                                                           603
      <210> 111
      <211> 595
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (595)
      <223> n = A,T,C or G
      <400> 111
```

graduation and a second con-

```
acatttaagt tcccatgtta cagaatccca tattgtgact atttcctcaa aactaactgc
                                                                                60
tagtaaagaa ccatcttcgg agaaacaaca gttagttgct tgatacttgt gataactacc
                                                                               120
aacaaagtca caggtccagc caacagcttt tttgtatatg tcagagtcat ctgttaatat
                                                                               180
ccatactttg aagtaaccat ctttgctagc tgtaaccaag gtgggctgtt cagatttttc
                                                                               240
tgcattacag aaacagagag ctgtaatgca gtcttcgtgt ggcatgttaa ttttagtgtt aagaataaac ccttgtgttt tcttattata catccacagt ttcatttgca attcaagctc
                                                                               300
                                                                               360
aagttteett ttettgeege tggteeactg gtgeaageea gttaceaaag eageeaatge aageettggt aagteaattt ggateagane ataateanta atatateetg etggataata
                                                                               420
                                                                               480
ctaaattgga tactggntat cactntggag agaataaact gcaggtggcn ggntttcatt
                                                                               540
caaaccaage tttagtettg gacaateatn aaccagngaa atacteetat ntttn
                                                                               595
       <210> 112
       <211> 523
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (523)
       <223> n = A, T, C \text{ or } G
       <400> 112
acaagagcta ttagagatgc tgccatatgg atgggcaaaa ctgagccaat cccacttagg
                                                                                60
aatggaagge ttggacatgg aagggaggat ataaacgagg agttggagaa aaacgcaagc
                                                                               120
ccagtttttg ctagagtgga aatgaaagtg ggaatgaggg tcttgtttt agtcctctaa
                                                                               180
ggaccaggaa gcaattttaa aacttccttg gtttttctga aagcagcata ttcaaaatgc
                                                                               240
cagcaaaaac tootaacaac tgcaaaacca aaagaggatc aaagctcacc aacatccctt
                                                                               300
cttattgctg aaaggctcta aaattcagga tgccctgttc ccttgtaaaa gggaaaataa
                                                                               360
ttaaagtetg atttatggta atcataccac atcacactte taaaaaaata ttteaagtgt
                                                                               420
gtgaccaggg gaccgtttga conccatttt attaaccttc actttantgg gaaaaataaa
                                                                               480
accttttcca gggccatttn atnccaggac ttttagtagg ggg
                                                                               523
      <210> 113
      <211> 578
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (578)
      <223> n = A, T, C \text{ or } G
      <400> 113
acagtgtaaa taactaagtt gttaactgtc aagtccagtt atgtattctg taagttgtgt
                                                                                60
tctagtcttt gactaaaatt tatcatctct tataatggga cttaatcttt ctctaaaagc
                                                                               120
atataagagc ttgtcaatag agcaatcaat caaaaagatt ttgtgattca taacattgaa
                                                                               180
gttagtctgg ttaagagttt tggtttagac ttcatttata ttttccttac taatatctaa
                                                                               240
tatttaatga ataatgatca attttttata aagttattaa tatgatcagg gaaacctttg
                                                                               300
ggacttctga caggcatctg gtgaagagac aattcaagcc ttagtgacta tttagaatag
                                                                               360
ccagtgatca ctagctaatt ctcatatcca tgcctttttt gccctggtta cagtcttaaa
                                                                              420
agaggtaaaa cagcaaatat tttttttaag ggaactataa ccctangaat tcctgaaaag aatttcaaaa aaaataagac cctgtggcca tggngnccaa acntaagacc tactatggct
                                                                              480
                                                                              540
atattggtcc attaaaaata aattactact aatccaaa
                                                                              578
```

```
<210> 114
       <211> 613
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (613)
      <223> n = A, T, C \text{ or } G
acggtagtaa gaaacctttg agatctttct gacttttcaa aattagagaa agcaaatqqq
                                                                            60
atggatagat ttttttttc ttttcaaggg gggcaggaag gtaatggttt gagtagcctt
                                                                           120
tgtttaaaaa aaaactaaat atatttaaaa ggccacattt atattttttt cacaagaacc
                                                                           180
acataataaa ttccacttct tgacctgaat ttggaaatcc gaaattacta atccaggcca
                                                                           240
gqtgtggtgg ctcatgcctg taatcccagc actttgagag gccgaggtgg gcagatcact
                                                                           300
tgaggcctgg agttcaagac caccttggcg aacacggtga aaccccgtct ctacqaaaaa
                                                                           360
aaaaanatat aaaaaaagta ctggttatta accaaccaqc ttaqaaaaat aatcatqqtn
                                                                           420
gacacnttan ttcattcttc taaaagcctg ttgatctggg ccttcctgtt gccagcattt
                                                                           480
cccctttttc aaaaatgggg ggccttttct ttaattnnac ctcgtggngn aananaattt gaagggcccc aggaagttnt ttgggcnctt tgaagcgttt tncacncgtn tagattctnt
                                                                           540
                                                                           600
gattaaatcc tcc
                                                                           613
      <210> 115
      <211> 190
      <212> DNA
      <213> Homo sapiens
      <400> 115
ggtacattgc cactgagtaa agagtggcac cagccacggt ggtaggtgga agaaacatag
                                                                            60
atcccaatga ggacacaaag acgagaccca ggcccactcc caggggtgca cccatgttca
                                                                           120
gaaacttttc actgggcgca cacatggcca cagtggagag gcctcccaca atgccagctg
                                                                           180
tqtacttttt
                                                                           190
      <210> 116
      <211> 610
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (610)
      <223> n = A, T, C or G
      <400> 116
ggtactcttg gtttatcaat gggacgttcc agcaatccac acaagagctc tttatcccca
                                                                           60
acatcactgt gaataatage ggatcetata tgtgccaage ceataaetea gecaetggee
                                                                           120
tcaataggac cacagtcacg atgatcacag tctctggaag tgctcctgtc ctctcagctg
                                                                           180
tggccaccgt cggcatcacg attggagtgc tggccagggt ggctctgata tagcagccct
                                                                           240
ggtgtatttt cgatatttca ggaagactgg cagattggac cagaccctga attettctag
                                                                           300
ctcctncaat cccattttat cccatggaac cactaaaaac aaggtctgct ctgctcctga
                                                                           360
gccctatatg ctggagatgg acaactcaat gaaaatttaa agggaaaacc cttangcctg
                                                                           420
```

```
aaggtgtgtg ccacttcaga gactttacct taacttgaga cngntcaaac ttgcaaacca
                                                                            480
 tggngnggaa atttgccgaa ctttacactt tgggcaggtt ttttcccaga agtcanaaca
                                                                            540
agaactectn ntettganaa gggttttane ecetttnaat ggeettgett atgetgeett
                                                                            600
 tttcgtttgg
                                                                            610
       <210> 117
       <211> 608
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (608)
       <223> n = A, T, C \text{ or } G
       <400> 117
ggtacgcggg gggtattatt tgtgccaacc aatgatgctt ttaagggaat gactagtgaa
                                                                            60
gaaaaagaaa ttctgatacg ggacaaaaat gctcttcaaa acatcattct ttatcacctg
                                                                           120
acaccaggag ttttcattgg aaaaggattt gaacctggtg ttactaacat tttaaagacc
                                                                           180
acacaaggaa gcaaaatctt tctgaaagaa gtaaatgata cacttctggt gaatgaattg
                                                                           240
aaatcaaaag aatctgacat catgacaaca aatggtgtaa ttcatgttgt agataaactc ctctatccag cagacacacc tgttggaaat gatcaactgc tggaaatact taataaatta
                                                                           300
atcaaatcat ccaaattaag titgticgtg gtagcaccit caaagaaaat ccccgtgact
                                                                           360
gctatagacc cacactaacc aaaggtcaaa attgaaaggt gacctgaatt cagactggat
                                                                           420
taaagaaagg tgaaaccatt actgaaagtg gatncatggg gaagccattt tttaaaaaat
                                                                           480
nccccaaanc attgatggga attccttnng gaaatacttg aaaggaaccn nnnnagacca
                                                                           540
                                                                           600
atcnttcc
                                                                           608
       <210> 118
      <211> 578
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (578)
      \langle 223 \rangle n = A,T,C or G
      <400> 118
actecaetta geaaatgeee tgeeageaaa gteacagatg aettttttae eeaatettag
gtaaatctgg attatctgcc caaccgtgca agtcaataag ccacccttga aaactgtgtc
                                                                            60
aagatttgag gaaacaggtc ttaagaacct atccaacaca tgattccata accaatacat
                                                                           120
cttangttgt tttaggcaaa taggtgtatc tcttgaatca ctgatggatt caatatcaag
                                                                           180
atctataatt ttcacgttta aaatttactc tgccgaggac attttattgg taaagcataa
                                                                           240
accagttagt ttgacagaca cnaaaaagaa aacnaaatgt tcacagtcct atcttcgtag
                                                                           300
ggattettgg ctataaaaat tggetteagg tteaaggtet tagaceacte ttetaagget
                                                                           360
nctactggat atantantta ccacttgggg nccaaactta aaacctcntg gactttttcc
                                                                           420
cettanggae nangaaaaac caaggggttg tggtttgaac teentacaet tggngnnaaa
                                                                           480
ncttttcttg gnngnatnta aanattaagg ggcttttn
                                                                           540
                                                                           578
      <210> 119
      <211> 584
      <212> DNA
```

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<213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (584)
      \langle 223 \rangle n = A,T,C or G
      <400> 119
actgtcttag aatattattt atttttttgt atttgtaaat ctgtggacaa aagagggttt
                                                                          60
cctcactcct tttactcact gggctcatga cagtgaagga gatgctccat ctgcttctcc
                                                                         120
cectttetet tgetgtagte caatgtgeta tgagcateag ettaetttge caettagage
                                                                         180
aagcaaaacc cagtgcaaga gtctcgttca gctctaaata ggtttgcttt cttttagtta
                                                                         240
cagtgeceat tttgaaattg cetatacagt ettagtgace atttaaaccg gacgaactan
                                                                         300
gcgtttaatt ttcacttctt catgttnaat tngcagttca anatttatag naagatggnt
                                                                         360
atttcgaaaa nacaaaaaan tggnttttta anaaaanaag tncnttggtc ggcgaancan
                                                                         420
gentaagggg egaattteea geneaactgg genggeeegt nnentagngg atceceaace
                                                                         480
tttggtaccc angettngge nntaancaat tggnecanag nttgttteec tggggtgaaa
                                                                         540
antingthatc ccgttcccaa ttcccnnaca ncnnaccnng cccq
                                                                         584
      <210> 120
      <211> 587
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(587)
      \langle 223 \rangle n = A,T,C or G
      <400> 120
acgcggggc cgtagcagcc gccgcccatc cctctttgtg tgctttggaa agccgcggag
                                                                          60
ctggtggtgg ctacagttgg tgttgggggc ttaggcgagg gacgttaccg ggaagttgca
                                                                         120
ggcgggagga ctcttcccca tccagtcacc tgacaggtca caaacatgtc agacaaaagt
                                                                         180
gaattaaagg ctgagttgga acgtaagaag cagcgactgg cccaaatcag agaggaaaag
                                                                         240
aagagaaaag aagangaagg gaaaaaaaa gaaacagacc anaataagga agctgttgct
                                                                         300
cctgtgcaag aagaatcaga tctttgaaaa aaaaaggaga gaagctnaaa gcatttgctt
                                                                         360
caaagcatgg ggctaacttc agaaatcccc ccattggncc ttcctnctaa tncttncatn
                                                                        420
ccttcaaaat ctgtggagcc ctttccaagg tgaaacttgn aannccaaga antntggaaa
                                                                        480
atggeneeet tggggaatet agacenaggg neetttttna acettggaat ngnttaaaaa
                                                                        540
tcacnccaag nttgactttt ccttccttcg anaaaattgg gtcccnn
                                                                        587
      <210> 121
      <211> 570
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(570)
      <223> n = A,T,C or G
      <400> 121
ggtactcttg gtttatcaat gggacgttcc agcaatccac acaagagctc tttatcccca
                                                                         60
```

```
acatcactgt gaataatage ggateetata tgtgeeaage ecataaetea geeactggee
                                                                           120
 tcaataggac cacagtcacg atgatcacag tctctggaag tgctcctgtc ctctcagctg
                                                                           180
 tggccaccgt cggcatcacg attggagtgc tggccagggt ggctctgata tagcagccct
                                                                           240
ggtgtatttt cgatatttca ggaagactgg cagattggac cagaccctga attettetag
                                                                           300
ctcctncaat cccattttat cccatggaac cactaanaac aaggtctgct ctgcttctga
                                                                           360
 agnectatat getggagatg gacaacttaa tgaaanattt aaanggggaa aaccettaag
                                                                           420
 cettgaggtg tgtgnecact teanaggaet ttaacettaa etttgagaee aggteaacet
                                                                           480
ggnaanccct tggtggagaa attggccgaa cttcccnact ttggccaggn ttttcccang
                                                                           540
 antgtcaaan caagacttcc ttatcatgnn
                                                                           570
       <210> 122
       <211> 551
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(551)
       \langle 223 \rangle n = A,T,C or G
       <400> 122
actateteta tteaggatta tgaagttttt egatgegaag atteaetgga tgaaagaaag
                                                                           60
ataaaagggg tcattgagct caggaagagc ttactgtctg ccttgagaac ttatgaacca
                                                                          120
tatggatccc tggttcaaca aatacgaatt ctgctgctgg gtccaattgg agctgggaag
                                                                          180
tccagctttt tcaactcagt gaggtctgtt ttccaagggc atgtaacgca tcaggctttg
                                                                          240
gtgggcacta atacaactgg gatatctgag aagtatagga catactctat tagagacggn
aaagatggca aatacctgcc cgtttattct gtgtgactca ctggggctga gtgagaaaga
                                                                          300
                                                                          360
aggeggnetg tgeagggatg acatatteta tatetttgae ggtaaceatt egtgatagat
                                                                          420
necagittaa ticecatgga atcaaatcaa attaaatcat catgactace tiggiteeee
                                                                          480
atcggttgaa gggacngnat tcattggggn ggcattggat ttgatnncna gntttattca
                                                                          540
atactttctc n
                                                                          551
       <210> 123
       <211> 575
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (575)
      <223> n = A,T,C or G
      <400> 123
acttaataca tattttcaaa cctgtttgca tttcaaacaa agttagcgtt tttgtaaatc
                                                                           60
aaatttgata accegactaa aaatatttte cagetttatt atttaaggag etgeacagee
                                                                          120
tttaaagtgg ggaccaggag gcaggcagag gcagagagac tgaatgcacc caggactgcg
                                                                          180
cagcagteta cagcaacatg teccacaact ttggtgetgg aaacacaagt aatgcacaag
                                                                          240
acagetgeec tecagtgtea ggatectgtg aaacageata teaaaagate gecagettet
tataatttac acactttcat ttaggattgc ttttttgaag aaaaatcttt aagaatgcca
                                                                          300
                                                                          360
tttttaattt aatatccaga accctggaat ttaaaaaaac ctaatngaaa ggaaattaac
                                                                          420
tggtaccatc aaaaatgggg ntgntggttg gancntgtgt gaagttaggg aattctatgg cttttttaa gatgccccgg aaaatttaac cccttaatng cangtttaat ttngaattcn
                                                                          480
                                                                          540
cnccaggtan tgtatgtnng gctcanatta gtanc
                                                                          575
```

```
<210> 124
       <211> 570
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (570)
      <223> n = A, T, C \text{ or } G
      <400> 124
actgagacaa tggttagggt tgttttctta attcttttcc tggtagggca acaagaacca
                                                                            60
tttccaatct agaggaaagc tccccagcat tgcttgctcc tgggcaaaca ttgctcttga
                                                                           120
qttaagtqac ctaattcccc tgggagacat acgcatcaac tgtggaggtc cgaggggatg
                                                                           180
agaagggata cccaccacct ttcaagggtc acaagctcac tctctgacaa gtcataatag
                                                                           240
ggacactgct tctatccctc caatggagag attctggnaa cctttgaaca gcccagagct
                                                                           300
tgcaanctag cettacecaa aangaetgga aangagaeat atetnteaag ettttteag
                                                                           360
gaangcgtnc ctgggaatcc aaggaacttt ttgatgctaa ttanaaangc ttgggactta
                                                                           420
aaaatgtccn ctanggngtg gcacttttac angtttttgg aangcttnga aggcaganng gggtcnaana ntnaaaanac nnttgacntg ntaatanngg aatantangg cnaatggaaa
                                                                           480
                                                                           540
ctgngttggg ggaggatcaa tttaaagagg
                                                                           570
      <210> 125
      <211> 593
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (593)
      \langle 223 \rangle n = A,T,C or G
      <400> 125
ggtacagaga tttaaatgaa atcttcgaaa gaataaattt gcttttcagt ccactgtatt
                                                                            60
ttcaaaattg attatcacca agettggatg aaagetgtga accacaaacc atttgtttat
                                                                           120
ttaatagaaa aaagaatgtg tagattatta gcaaagtaat gccttaaaat gtatcttcac
                                                                           180
acagttgaaa ttttagtata aacttgtata tcaagttgct ttccattatt tattctactt
                                                                           240
taaaaatata tacaactatg atgttcaaat atgtattctg agccattatg ttcaaacata
                                                                           300
aatatctggg aaattcaaac tgctgcaaca agttaggaaa ggattaagga aaaatgatga
                                                                           360
gctacaaatt atgtagttgg aggaagaaaa aaatgttact tagcatttat gtctggatag
                                                                           420
gtatgtattt totaatttac atacacatat ccagttgagt atagaccacc atcaaaatgt
                                                                           480
accagttaca cagagactag actaaaccac cctatttcta tacaggtacc atagtggatt
                                                                           540
caaaaattta atatctcata gttcccaaaa ttattgnggn aatatgctna ttt
                                                                           593
      <210> 126
      <211> 592
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(592)
```

```
<223> n = A, T, C or G
       <400> 126
acgcgggggg gccttccggg acgagggcgc gtgggtgagg aaggtcaggt ctaggaactc
                                                                              60
taacteettg ccacteaaga aatgteetee ettteagaat atgeetteeg catgtetegt
                                                                             120
ctcagtgccc ggctatttgg tgaagtcacc aggcctacta attccaagtc tatgaaaqtq
                                                                             180
gtgaaactgt ttagtgaact gcccttggcc aagaagaagg agacttatga ttggtatcca
                                                                             240
aatcaccaca cttacgctga actcatgcag acgctccgat ttcttggact ctacagagat
                                                                             300
gagcatcagg attttatgga tgagcaaaaa cgactaaaga agcttcgtgg aaaggagaaa
                                                                             360
ccaaagaaag gagaagggaa aagagcagca aaaaggaaat agtgttggtc ccttcaagag
                                                                             420
ggagactttc ttcctaatgg ccggaaagaa gaaagtgcat ttattgqctt tccacatatt
                                                                             480
ggaggaatgt catcttccta aatgaagttt atttggagga acacagtcat ttccttqqtq
                                                                             540
aaactaatcc ggtacattgn ggttgggttt ttgaacacat ctactgggca aa
                                                                             592
       <210> 127
       <211> 600
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(600)
      \langle 223 \rangle n = A,T,C or G
      <400> 127
acagtggtcc ttttcagagt tggacttcta gactcacctq ttctcactcc ctqttttaat
                                                                              60
tcaacccagc catgcaatgc caaataatag aattgctccc taccagctga acagggagga
                                                                             120
gtctgtgcag tttctgacac ttgttgttga acatggctaa atacaatggg tatcgctgag
                                                                             180
actaagttgt agaaattaac aaatgtgctg cttggttaaa atggctacac tcatctgact
                                                                             240
cattettat tetatttag tiggttigta tettgeetaa ggtgegtagt ecaactettg gtattaceet ectaatagte atactagtag teatactee tiggtgtagtg tattetetaa aaagetttaa atgtetgeat tigeaneeage cateaaatag tigaatggget etetttigge
                                                                             300
                                                                             360
                                                                             420
ntggaattcc aaaacntcag agaaatggtg tcatcaagga gaaccttcat aaccccntga
                                                                             480
anggattaaa aagccccaaa tggggggaac tgataatagc acttaaggct ttaagaattg
                                                                             540
gncacanttt caccttgtga acccattnna cnatngngcc taanngctnc ctnctncaan
                                                                             600
       <210> 128
       <211> 588
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc feature
       <222> (1) ... (588)
      <223> n = A,T,C \text{ or } G
      <400> 128
ggtacttttt ttttttttt ttttttttt tttttttgag acggagtctc actctgtcac
                                                                              60
ccaggctgga gtgcagtggc atgatcttgg ctcactgcaa gctctgcctc ctgggttcac
                                                                             120
gccattetee tgeeteagee teetgagtag etgggaetae aggegteege caccaegeee
                                                                             180
agctaatttt ttgtattttt ggtananaca gggtttcacc gngttagcca ggatggnete
                                                                             240
catctcctga cctcgtgatc tgcccacctn ggccttccaa agtgctggga ttacaggcat
                                                                             300
gagecaegge geetggeeag gatggtatat tittaactee ticactggge eccaeceetg
                                                                             360
```

```
actiticizet tiangaggie tggggtgagg etgaanatet gggggeeaca etteqagage
                                                                        420
aaccaagact gtaagtgggg ccttccanag cccaatgaag ggaatactta ggtacaggan
                                                                        480
gtgtctgcat ggncncangt gtggggtttn cttctcggcc ttaaccaqaa aqtatctctq
                                                                        540
gttttaattt taaaatqaaa attttaaaqq qtqnctqaaa cnaattqq
                                                                        588
      <210> 129
      <211> 588
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(588)
      <223> n = A, T, C \text{ or } G
      <400> 129
ggtactgccc tctccagatc agcagttcag gagagcacag gaggcaaaac acagattgct
                                                                        60
gggcttattg gtgccatcat cgtgctgatg gtcgttctag ccattggatt tctcctggcg
                                                                        120
cetetacaaa agteegteet ggeagettta geattgggaa aettaaaggg aatgetgatg
                                                                        180
cagtttgctg aaataggcag attgtggcga aaggacaaat atgattgttt aatttggatc
                                                                        240
atgaccttca tetteaceat tgteetggga etegggttag geetggeage tagtgtggea
                                                                        300
tttcaactgc taaccatcgt gttcaggacc caatttccaa aatgcagcac gctggctaat
                                                                        360
attggaagaa ccaacatcta taagaataaa aaagattatt atgatatgta tgagccagaa
                                                                        420
ggagtgaaaa ttttcagatg tccatctcct atctactttg caaacattgg tttctttagg
                                                                        480
eggaacttat egatgetgnt ggetttagte cettegaatt taegeaageg cacaaacttt
                                                                        540
gaggaaaatc cgaaactgcn aagcaagntt gntacaagtg acccaaan
                                                                       588
      <210> 130
      <211> 190
      <212> DNA
      <213> Homo sapiens
      <400> 130
ggtacaaaaa aaaccttaca taaattaaga atgaatacat ttacaggcgt aaatgcaaac
                                                                        60
cgcttccaat tcaaagcaag taacagccca cggtgttctg gccaaagaca tcagctaaga
                                                                       120
aaggaaactg ggtcctacgg cttggacttt ccaaccctga cagacccgca agacccgcg
                                                                       180
tactttttt
                                                                       190
      <210> 131
      <211> 386
      <212> DNA
      <213> Homo sapiens
      <400> 131
ggtacagaac tcagaggaaa aaagaaatta aattttagct ttctggagag cagccctct
                                                                        60
ctggcaccat caaacacttc tttgtttccc ttcaacttgg aactcttcaa acatcagggg
                                                                       120
ttgtgagggt ttggccattc ttttatcttg ggtccatgtg agtgacagaa atggtgcggc
                                                                       180
ctgggaaaga tctccctcct ttacattttc tcttctcct cctcctctt attctaaaac
                                                                       240
tgtgcctcca acagagggc aggggctctt gtagagagat ccctggccca ggacaggaga
                                                                       300
tgccaaatct aatttatctc actgagggcc tttgagaaaa acgcttcagg gccaggctca
                                                                       360
gtggctcatg cctatataat cccaqt
                                                                       386
```

<210> 132

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<211> 593
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(593)
      <223> n = A, T, C \text{ or } G
      <400> 132
actgagacaa tggttagggt tgttttctta attcttttcc tggtagggca acaagaacca
                                                                         60
tttccaatct agaggaaagc tccccagcat tgcttgctcc tgggcaaaca ttgctcttga
                                                                         120
gttaagtgac ctaattcccc tgggagacat acgcatcaac tgtggaggtc cgaggggatg
                                                                         180
agaagggata cccaccact ttcaagggtc acaagctcac tctctgacaa gtcagaatag
                                                                         240
ggacactgct tetatecete caatggagag attetggcaa cetttgaaca geecagaget
                                                                         300
tgcaacctag cetcacccaa gaagactgga aagagacata tetetcaget ttttcaggag
                                                                         360
gegtgeetgg gaateeagga actittigat getaattaga aggeetggae taaaaatgte
                                                                         420
actatngggt gcactctaca gtttttgaaa tgctaggang cagaagggca aaaataaaaa
                                                                         480
acatgacctg gttgaaggaa naaaagcaaa gaaacttggg ngggaggaca attaaaaaga
                                                                         540
gnncctggga teceetntte ttaggteeet etettaenaa ggaenetntt tat
                                                                         593
      <210> 133
      <211> 588
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (588)
      <223> n = A, T, C \text{ or } G
      <400> 133
acagancatt nnnagenetn geacaggnta cagaacetna cagacecaaa ggaacategg
                                                                         60
ataggenaag egactaeagg aggegtgtgt gegettggge naggtaaaca gggteagtat
                                                                        120
tggtcnngtg acaagagnca cgaantctgg ccngacantg angtnaanaa ggttnatnnt
                                                                        180
ttnacantta tnnnanatat nnnnnaannt attaanctgc ancanntgat tttnacacct
                                                                        240
anttactaga aaactaanga aagcactnat tagctctgaa tnaantnaca tggnaagcct
                                                                        300
tttactaatc tncaaanaaa ccttctctgc antatnnnaa agattttatn atacaangng
                                                                        360
gnnnatcnct cnatcatann gggttctatt ananaaccct gctaantntg cgacttacag
                                                                        420
aacanccage ntananatga ntttcatgee catttgggaa geatngeeg ggtatcacaa
                                                                        480
aggaaaccta ctaaagnttt ctgttatacc agccttcntt cntatcantg catgngnana
                                                                        540
nanaacentt gaaggttnte enggggaett tnttetnttn etttgeee
                                                                        588
      <210> 134
      <211> 618
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(618)
      <223> n = A,T,C \text{ or } G
```

```
<400> 134
tenageggee nneenggeag gtacanteae annttnnang anetnaacae anactanetg
                                                                                 60
nngtcaaata ttnaacaaaa gcantagatg aanctgctta acattcacgg aaaaacaacc
                                                                                120
aaaagaaggg aggggtgata aaccanaaaa atgantgacn aaaactaaga gacctcatan
                                                                                180
gngtetttae aatenggaat teagatgeaa ggaacagaen caaanetgte taaaatgtna
                                                                                240
cctatgaggc nacanaaagt gacttaaagt ctggtntnan taaaaaatga caacccttat cctagagagt cttacnttat ttaatccana cnttatntaa cgccncngat ttttgnttgg
                                                                                300
                                                                                360
ngctatggng ttnattttnt atcagaanga antgtgggac anatgcatta ctgnttgttn
                                                                                420
aaagngcttn acagetaatt cacnecenng ggcatggtca aaaaggnaan aacenggnca
                                                                                480
tatattgntg anatgaaaaa accacntgtt aaaaaaataa ntgnagccna ntgngttttn
                                                                                540
natgataacc aaatnttnac nttcagtann ngccttttan aagttggtga actccgaaat
                                                                                600
ctncttttt aaaccngg
                                                                                618
       <210> 135
       <211> 374
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(374)
       \langle 223 \rangle n = A,T,C or G
       <400> 135
acttttttt ttttttttt ttttttttg gggatggagt ctcactctgt tgtccaggtt
                                                                                 60
ggagtgcagt ggtgtgatct cggctcactg caaccintgc ctcccaagtg attctcctgg
                                                                                120
ctcancetee tgagtagetg ggactacagg catgeactae catgeeegge taatttttgt atttttagta nanacagggt tteaceatgt tggeeagget ggtettgate tectaatete aggtgateeg cetgeetean cetectaaag tgetgggatt acaggeatga gecaetgtgt
                                                                               180
                                                                                240
                                                                                300
ntggccaana ncactcgtaa gaaggatggc agtatcacaa aatcaagcca gagatacaga
                                                                                360
gattacccgc gtcc
                                                                               374
       <210> 136
       <211> 581
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (581)
      <223> n = A, T, C \text{ or } G
      <400> 136
actccagcct tgctgaagct gcctcaaagg ctgatggttt ggcagttatt ggtgttttga
                                                                                60
tgaaggttgg tgaggccaac ccaaagctgc agaaagtact tgatgccctc caagcaatta
                                                                               120
aaaccaaggg caaacgagcc ccattcacaa attttgaccc ctctactctc cttccttcat
                                                                               180
ccctggattt ctggacctac cctggctctc tgactcatcc tcctctttat gagagtgtaa
                                                                               240
cttggatcat ctgtaaggag agcatcagtg tcagctcaga gcagctggca caattccgca
                                                                               300
gccttctatc aaatgttgaa ggtgataacg ctgtccccat gcagcacaac aaccgccaac
                                                                               360
ccaacctctg aagggcagaa caagtgagag cttcattttg atgattctga gaagaaactt
                                                                               420
gtncttctca agaacacaac cctgcttctg acataatnca ataaaataat aattttaaaa
                                                                               480
aataaattat ttcaatatta ncaagacaca tgccttnaat natctgtaaa ctaaaaacta
                                                                               540
aaatttantc tactgnttaa tcnaanataa taatagcttc a
                                                                               581
```

```
<210> 137
       <211> 504
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (504)
       <223> n = A, T, C \text{ or } G
       <400> 137
ttttncaaan nnaagttttt tacttccnaa aantnatggc taaggggngg gnggngggng
                                                                             60
aaaaaagnaa aacaaaaaaa ccccaaaaaa atggggnggn naaaaggggg gganaaaaaa
                                                                            120
conntntttt ntaaantntn acaaggcaag ngcnnangga aaaaaaaan ncctgnaaaa
                                                                            180
tececenegg nnggggnaaa natnnnggtt teettttgnt ttnaaaeeen ntnangnaag
                                                                            240
gntntccccc ntnccctna atnaaaaatt tntntnccng ggccnnaacc ncentanggg
                                                                            300
naaattccac cncnctgggg gccgttanta agggatccna gctnggccca ancttggnga aacatggcaa aactgttcct nnggnaaaat gtttcccctc anaattccca naaaataaaa
                                                                            360
                                                                            420
ceggaacata aagngaaaac engggggeet aagngggnen cacnecattt attggggtgg
                                                                            480
cccncgnccc tttcaaangg aaac
                                                                            504
       <210> 138
       <211> 386
       <212> DNA
      <213> Homo sapiens
      <400> 138
acaacaaata acactgtgac tccaacctca caacctgtgc gaaagtctac ctttgatgca
gccagtttca ttggaggaat tgtcctggtc ttgggtgtgc aggctgtaat tttctttctt
                                                                            60
                                                                           120
tataaattct gcaaatctaa agaacgaaat taccacactc tgtaaacaga cccattgaat
                                                                           180
taataaggac tggtgattca tttgtgtaac tcactgaagc caaaatacta tcttttaaga
                                                                           240
tgtcccacat ggaagacgct attccaggat ctttaaattt ccatggatgc atataggatg
                                                                           300
tttgggagca tcatccgtga agaaaaaatc aattaaatca ttgtgttcaa caggaatatt
                                                                           360
taaaataaaa aaaaaaaaa agtacc
                                                                           386
      <210> 139
      <211> 586
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (586)
<223> n = A,T,C or G
      <400> 139
ggtactcaag tttataatgt ccccaaacct taagactaga aaatcatccc aagaaaaagg
                                                                            60
cctatagttg gtttaatttc accctgagaa tactgtgata aaaatcaata tatttcagag
                                                                           120
ctagtaagta tttaaaaatt agtgtctcaa aaaggggaca tcataaggga aatacagggt
                                                                           180
ttagaggtet gageteaagt ggtgtaagae agttettet tetteeteet ttaaactett
                                                                           240
cactttgctc taacacggaa gatgggggac agtgatcccg aaggtattac taaaatattg
                                                                           300
cagettteag taattatgag aageacagat ateaccagaa aagaaageaa teatttggag
                                                                           360
```

```
tactaagaaa cgaaacaatg ttatttggtg gtgtataatt ctacttttct agtagattac
                                                                         420
tgngtggaat tctgtgaaaa atatttgaga aaangcctgt attgcataaa taaatctttg
                                                                         480
tatgttgcaa aaaaaaaaa aaaaaaaagt acctgccggc cgncccaang gcgaattcca
                                                                         540
cacctgcggc cgtctagngg tccacccggt ccacttgggt atatgg
                                                                         586
      <210> 140
      <211> 591
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (591)
      <223> n = A, T, C \text{ or } G
      <400> 140
acagggagga atttgaagta gatagaaacc gacctggatt actccggtct gaactcagat
                                                                         60
cacgtaggac tttaatcgtt gaacaaacga acctttaata gcggctgcac catcgggatg
                                                                        120
teeetgnace aacetteaag geenaaacee nnntggtgnn tttggnetnt aaatnaggat
                                                                        180
ggccctgtnt tccntaggta acttgttccg ttggtcaagt tattggatca attgagtata
                                                                        240
gtagtteget ttgactggtg aagtettnac enngteentt tngngtgggg tttttttagg
                                                                        300
naaaagnett ttggtneatt nntggggggg gnaggggaet gaacetttat tntttecaaa tneaeettaa anteagggae aanaaacatt ceaanaacea caatettta aaaaattaae
                                                                        360
                                                                        420
tngccagtgg gaatgtttaa aaanntnaan ggtctttttt gccttggttt ttgtgggggt
                                                                        480
ctctcttccc ccccctgggg ttaatttttn aagccgggac ctcncnaana ccccttttt
                                                                        540
591
      <210> 141
      <211> 592
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
     <222> (1) ... (592)
      <223> n = A, T, C \text{ or } G
      <400> 141
ggtacacaaa ccaagacaat atcagggtga caggtgaatg aacttaaatt ctcagtcttg
                                                                         60
totattcacc aaaaaagtat actgcctgtt ttttctttaa ttattcaagg ttgatgactt
                                                                        120
ttaggaacat gttttatact gtatttttta attaaagcaa gtgccttgat gtaattccat
                                                                        180
gtaaatcatt gcttaaccct cttatgggat gaggatgagt tattaatgta ttgcagccta
                                                                        240
ctggaaagga gggggagttg gttaatagca gatacttttc ttctagaagc ttatgtttta
                                                                        300
tgctgtttat tatgtaagat cctgtatgtg tgttgagatt tagaggtttc atttgttttg
                                                                        360
tctgctaata aattgttact ctaataataa ccnngnnaaa naaannnnnn nnnnnnnnn
                                                                        420
nnnannnggt neetgeeeng geggeegete gaaagggega attecaneea etggenggeg
                                                                        480
gtactaaggg gatccgnctc gggncccaac ttggcgtaat atnggcatac tggttcccgg
                                                                        540
gngaaatggt atncgtcaaa ttccccaaat acnaccggaa ncttaagggt aa
                                                                        592
      <210> 142
      <211> 595
      <212> DNA
      <213> Homo sapiens
```

```
<220>
       <221> misc_feature
       <222> (1) ... (595)
       \langle 223 \rangle n = A,T,C or G
       <400> 142
acaacacctt cattettaat gettettagg geateacagg ttttagaaat taatgtattt
                                                                             60
ttagcattcc acagtaatga tcactttcaa aaactgcaat atacatctgc atgttacact
                                                                            120
gacatacaac acataagtat tttgtcacac atcaactttt agcctcaaat aatagaatac
                                                                            180
aaaaagctac actggacata acaccaccga acttttgaat atcccctttt cccaattgtt
                                                                            240
aacaggtagt actgggatta caggcgtgag cctctgcgcc tggccaagtg gaggttatta
                                                                            300
ttaaccctat ttaacagata taaaaagaag agattagaga attttatcaa tgttcccact
                                                                            360
gtcaaataga atataagcaa tgatacaaaa tgttgagtct tcatcctcta actccagatc
                                                                            420
ctggtatatt gccctacatt tctatacatt aatactaact tatacactga atacaagagt
                                                                            480
naaaccaact gtengggett aatanggnga aaatgetett gneetaaane accagggtgg
                                                                            540
ctnggtttat tcctacatgt ggactaaaan gnaatcatct ttatggengg aaana
                                                                            595
       <210> 143
       <211> 620
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
       <222> (1) ... (620)
       <223> n = A, T, C or G
       <400> 143
actactegat tgtcaacgtc aaggagtcgc aggtcgcctg gttctaggaa taatggggga
                                                                            60
agtatgtagg agttgaagat tagtccgccg tagtcggtgt actcgtgtga agttggcagg
                                                                            120
gacggtteet gteatettet tgggettatt tggtgtgetg ttgaagggg gagactagag
                                                                            180
aaatggcagg gaacctctta tccggggcag gtaggcgcct gtgggactgg gtgcctctgg
cgtgcagaag cttctctctt ggtgtgccta gattgatcgg tataaggctc actctcccgc
                                                                           240
                                                                           300
cccccaaagt ggttgatcgt tggaacgaaa aaagggccat gttcggagtg tatgacaaca tcgggatcct gggaaacttt gaaaagcacc ccaaagaact gatcangggg cccatatgct
                                                                           360
                                                                           420
tcgaggntgg aaanggaatg aattgcaacg ttgtattccn aaagaagaaa atggttggaa
                                                                           480
gtaaaatgtt cettatgace tencaacett ataaacneat cegittnitt acaacentta
                                                                           540
accacatggg aagttcattn aaaaaactg aaaactttgn aaagnttttt ttnnccttga
                                                                           600
aaagggaact tacctcqccc
                                                                           620
      <210> 144
      <211> 613
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (613)
      \langle 223 \rangle n = A,T,C or G
      <400> 144
cgaggtactt ttttttttt ttttttttt ggggtcagtg gtgatatccc cctaatcaat
                                                                            60
```

```
totgattgng ttoottttaa tottototoa tttottttt attagactag atagtgattt
                                                                                 120
atctatttta ttaatttttt caaaaaatca cctcctanat ttgttgtttt ttaaggggtt
                                                                                 180
ttatgtetet ateteettea gtteaactet gatettggnt atttettgne ttetgetaga
                                                                                 240
tttggggttt gntttctgnt ggntctctaa gttctttttg ntgngacatt agattgncaa
                                                                                 300
cttaaaatct ttctagctat ttgacgtggg catttaatgc tataaatttc ctggtaacac
                                                                                 360
tgctttcgct gtatnccana naatctggga tggtggggcc ttggtttcaa taanttccaa
                                                                                 420
tacctcttaa gggggnggag ccaanaagan ctaatagggg cagcactgct ctgggctncc
                                                                                 480
atcaanaagg acaaaaactg ggagngaccc tgcttnttca ctgaggnacc ggcccggccg
                                                                                 540
gccgtccnaa ggcgaatcca cncnctggcg gccgtctatg gatccacccg gnccaactgg
                                                                                 600
ggaatatggc aaa
                                                                                 613
       <210> 145
       <211> 345
       <212> DNA
       <213> Homo sapiens
       <400> 145
acactgatet acaaaaattt taaaatgage egggegeggt gaeteaegee tgtaateeea
                                                                                  60
gcactttggg aggccaaagc aggcggatca tgaggtcagg agatcaagac catcctggct
                                                                                 120
aacacggtga aaccccgtct ctactaaaaa tacaaaaaat tagccgggtg tggtggcggg
                                                                                180
cacctgtagt cccagctact cgggaggctg aggcaggaga atggcgtgaa gccgggaggt ggagcttgca gtgagccgag atcacaccac tgcactccag cctgggcaac aaagcaagac
                                                                                240
                                                                                300
tetcaaaaaa gaaaaaaatt tttttttaaa tgagetgggt gtace
                                                                                345
       <210> 146
       <211> 475
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(475)
       <223> n = A, T, C or G
       <400> 146
actacaaggt ttagcatttg ctctgctggt cgacattccc ccagtctatg ggttgtatgc
                                                                                 60
atcetttte ccagecataa tetacetttt etteggeact tecagacaca tateegtggg
                                                                                120
tccgtttccg attctgagta tgatggtggg actagcagtt tcaggagcag tttcaaaagc
                                                                                180
agteccagat egeaatgeaa etaetttggg attgeetaae aactegaata attetteaet
                                                                                240
actggatgac gagagggtga gggtggcggc ggcggcatca gtcacagtgc tttctggaat catccagttg gcttttggga ttctgcggat tggatttgta gtgatatacc tgtctgagtt cctcatcagt ggcttcacta ctgctgctgc tgncatgttt tggtttccca actcaaattc
                                                                                300
                                                                                360
                                                                                420
atttttcaat tgacagtccc gtcacacact gatccagttt caattttaaa agacc
                                                                                475
      <210> 147
      <211> 629
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (629)
      <223> n = A, T, C \text{ or } G
```

```
<400> 147
cgaggtacgc gggatttgaa tcttaaactg tatttttctc ttagtattgc taatgagtaa
agaaaagtct cataaggtag ccaaatgaaa aagaatgaaa gggaaagtga aaaattaagg
                                                                         60
                                                                        120
ggacaaaaga tgggatgtga aaagaagaat tctagtttga tggtgactca tattcacgat
                                                                        180
aggatacaaa gigigaitig tiggaaacat gicccaaait totaaaatto igctictoig
                                                                        240
ccaaaagcaa tgtctttctt ggttgatatt tgagttttaa aagggtcaaa tctttctaat
                                                                        300
tttttgtatc tttagagggc agcactagaa gaaatcagca ggtctaatcc caccagtaag
                                                                        360
aaaactacca cttcttgatt tttacagatt taaaaaaatc ttttcagtgc ctttctttt
                                                                        420
aatgtaaata caaatttaaa cctangctta atatagcgtt tccctttccc caagtgatgt
                                                                        480
cnaggtegat gecaaateaa tgateenaaa tgategnggt naaaataaet caaagggtte
                                                                        540
ttaaggngag tngcatgcca aaaaatacct tgattccggg ggtttggacc tggctttgtt
                                                                        600
ggggcctntg aaatgccaan ttancccan
                                                                        629
      <210> 148
      <211> 614
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (614)
      <223> n = A,T,C or G
      <400> 148
acaaaagagc ctgattcttt ttaattccac aaatacctag catctcaaag taacatgtaa
                                                                        60
acaaacttct atgctgctca atgaatcctt ccaatttcga taataaacta aatagtattg
gatctagtat atgactttca tgtgtaagtt atggttctat ccattacttt aacaatatta
                                                                       120
ctgatgtaac agagaaaaat tttcaactat tgtatttatt taaaacaaac tgacaagttc
                                                                       180
                                                                       240
aagcacctgt cttcagaaaa gccagcagca ttttttttt ttaacatact caaagtaaga
                                                                       300
tttggcctaa gcccttaata cctttctgaa cagccatgca actaaacacc ctcagggaga
                                                                       360
tgttacataa gggagagaag aacatggagc aatttgcact ttttccctag ataatattaa
caaggnaaag caaatncaga tctttatgaa tgaatggntg gcatggttaa tcacttggac
                                                                       420
                                                                       480
tttttaaact agagneneta teatattggt aaatagaaan aaaggatttt aataaagete
                                                                       540
tnectgette aaaattaagg ggaenttite tgggaggett teagggaeea taataaggta
                                                                       600
aaaggggacg gttg
                                                                       614
      <210> 149
      <211> 628
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (628)
      <223> n = A,T,C or G
      <400> 149
nccgaggnac tttnnttttt tttttttt ttttnaacag cgncttttca tttttattac
tcaaaaaagt ttcatttttt tatttaagct ttctgactct gngcttgggc cttcaacact
                                                                        60
                                                                       120
ttcacaacga ttttctgctc ctcgataagg aaagcccgct tgatcctana aaggaaaata
                                                                       180
ccaaattaat catttctta aaatgaactt catttttat ttagcccaaa aaaggnaaac
                                                                       240
atggtaaaga accaagcnaa gcaatcaggg aacccaggaa actacnggat acccaaatac
                                                                       300
```

```
ngagtaaaac ttaaaagggg aaattcattt aaagcaggga aatccctcaa tttcatgccn
                                                                              360
 gtagttatct gncctcctct gagcaagaat aactatgaag catcccccag gagaccacnt
                                                                              420
 atgagaetta attattggta ggatecagga atagnggnat ttnttgattt gcaaaangtn
                                                                              480
 taaaaaattt taaccctntt ttgaaaattc ccagnaaaaa caccncataa ggggctntgt
                                                                              540
 gttaaaacta aaattaaagg gaagggtttt tccagaaacc cccccanac cagggtttna
                                                                              600
 accggttang gcanntcncc aaaccnan
                                                                              628
       <210> 150
       <211> 509
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (509)
       \langle 223 \rangle n = A,T,C or G
       <400> 150
ttggggaann aaaaaaaac ttttttttt nggggnnngg ggntgcnanc natncaaaaa
                                                                               60
tcaaaancnt ntttgggttt taactttttt ttttttgntt gncaaannaa aantaaantt
                                                                              120
tntttttana tttgctaang ggccngancn gcnnaaaaaa nccttttttn ggggaanctt
                                                                              180
nggggcaaat tnnttnanen accetttggg anaacttttn ttaggggggn nnnaaccgne
                                                                              240
attititgece actitititee ettitignita anggggneet tgggenggae enceettagg
                                                                              300
ggnaattcac cenetggggg gegttatntt ggatecacte ggnecaactt gggggaaaaa gggaaaacnt tttetggggn aaatttttte cenenaaatt eecaanaana aaaceggaac
                                                                              360
                                                                              420
nnaaanttaa acceggggge ccaaggnggg cennecentt nttgggtggg ccetgecent
                                                                              480
ttaangggaa attttggccc tttttaaaa
                                                                             509
       <210> 151
       <211> 622
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(622)
       <223> n = A,T,C \text{ or } G
      <400> 151
ggtacttttt ttttttttt tttttttgc tttggacaaa tttattgaaa catacaggcg
                                                                              60
getgttagea gagaaateat tecatgattg atgtgttaca tttggceaet acettgaatg tataaatttaa aaattatatt tttcacaaet aageetttgg ccaaaaaagt catttageae
                                                                             120
                                                                             180
atctttaaag atcaataaga aatggatttt ggacattaaa aagatcaagt cactgaatta
aacagtagca accccatta atctagaatc ccatagtgct gaaggtagag gtgtctgtgc
                                                                             240
                                                                             300
aaagctagtc atttgttaac agcaatcana aaanatgggg gcaggcacac ctgtcaaaag
                                                                             360
tggcaacana nctggcagga caggacggct gggctggtct ggtcaggtga gcatgtncca
                                                                             420
aaaacagcag caacagaaaa cccgtccacc angcttgtga agcangtgga tggtcctagc
                                                                             480
tcatctnttn ttttggnctt ntancacata cactgngggt ttangangnt tctgaggncc
                                                                             540
accttgccnc cctacctgcc cgggnggccg ttnaaagggg aattccacca ctgggggccg
                                                                             600
tctaatggga cccacctggg cc
                                                                             622
      <210> 152
      <211> 313
```

```
<212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (313)
      <223> n = A, T, C or G
      <400> 152
acggtggatt agttcttttc agcatgttcc ttctgtatga tacccagaaa gtaatcaagc
                                                                      60
gtgcagaagt atcaccaatg tatggagttc aaaaatatga tcccattaac tcgatgctga
                                                                     120
gtatctacat ggatacatta aatatattta tgcgagttgc aactatgctg gcaactggag
                                                                     180
gcaacagaaa gaaatgaagt gactcagctt ctgcttctc tgctacatca aatatcttgt
                                                                     240
300
aaaaaaagt acc
                                                                     313
      <210> 153
      <211> 620
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (620)
      <223> n = A, T, C \text{ or } G
      <400> 153
cgaggtacgc gggagggcaa caagaaccat ttccaatcta gaggaaagct ccccagcatt
                                                                      60
gettgeteet gggcaaacat tgetettgag ttaagtgace taatteeet gggagacata
                                                                     120
cgcatcaact gtggaggtcc gaggggatga gaagggatac ccaccacctt tcaagggtca
                                                                     180
caageteaet etetgacaag teagaatagg gacaetgett etateeetee aatggagaga
                                                                     240
ttctggcaac ctttgaacag cccagagctt gcaacctagc ctcacccaag aagactggaa
                                                                     300
agagacatat ctctcagctt tttcaggagg cgtgcctggg aatccaggaa ctttttgatg
                                                                     360
ctaattagaa ggcctggact aaaaatgtcc actatggggt gcactctaca gtttttgaaa
                                                                     420
tgctaggagg caaaaggggc agagagtaaa aaacatgacc tggtagaagg aanaaagcaa
                                                                     480
aggaaactgg tggggaggat caattagaga ngaggccctg ggatccncnt nttcntaggn
                                                                     540
conteteata enaaggacae tttttatatg cotteecaaa etgntnggga agggtnaaae
                                                                     600
caaaatccgg ggtanaacct
                                                                     620
      <210> 154
      <211> 339
      <212> DNA
      <213> Homo sapiens
ggtacctgga ggatatagac ctgaaaacac tggagaagga accaaggact ttcaaagcaa
                                                                      60
aggagetatg ggaaaaaat ggagetgtga ttatggeegt geggaggeea ggetgtttee
                                                                     120
tetgtegaga ggaagetgeg gatetgteet eeetgaaaag catgttggae cagetgggee
                                                                     180
gteceetet atgeagtggt aaaggageac ateaggactg aagtgaagga tttecageet
                                                                     240
tatttcaaag gagaaatett etggatgaaa agaaaaagtt etatggteea caaaggegga
                                                                     300
agatgatgtt tatgggattt atccgtctgg gagtgtggt
                                                                     339
```

<210> 155

```
<211> 450
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (450)
      <223> n = A,T,C or G
      <400> 155
cgaggtactt ttttttttt tttttttt tttttcntat ttttgtttaa tttatttaan
                                                                           60
accacctnct tacaacttnc anagagaaaa tacaaaacaa gaaacanact tggtttnaaa
                                                                          120
tgcataacca gntgctggan tttaaagcat tactgataac attgttacan aanaatggca
                                                                          180
nnttactona gggcactina gtattoctna ggaataaaca ttgatttoto ttgtoctoo
                                                                          240
nntgggatgt tctcangtna agtcactgcn cctgcnctta gacatatttt ccatgtnnca
                                                                          300
naananggag cctgnaaant atgctnacag tnggaataag ccattnctaa ttccatqcca
                                                                          360
naaccnangg ctaatggnnc attettttt aataaggtat gtggaaaana ttentateee
                                                                          420
aaanaaaant tgcccggncg gtctntntaa
                                                                          450
      <210> 156
      <211> 760
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (760)
      <223> n = A, T, C or G
      <400> 156
cgaggtactg ccccagtgaa aatggaactg aaagagcctg tagctgtcag agaaaggacc
acetticage actgateggt tategtigte etcaaaatti acatggaagg aatgeeceae
                                                                          120
attgataatt tetttggetg tggetgggtt gtaggagaca ctaataggtt teagagaggt
                                                                          180
gtcatgtttg gtttcactgg ttttaatatc aacaggggac tggttatttc cattggcaat
                                                                          240
gggatacage ttgetecatt gttcaggace atttttgtca teatatecee agtetqqaet
                                                                          300
tgccattatc ttctactgag ttttcttttt ctgaaaacaa aaataatacc tggaataact
                                                                          360
aactgccccc gcgtcctgcc cgggcggcca aaggggcaat tccaccactg gcggccgtac
                                                                          420
ttatggatcc aactcgtccc ancttggcgt aatatggcat aactgttctg nggnaaatgt
                                                                          480
atcccttaca attcccncac atcnacccga acctaantgt aancctnggn gcnnataagg actactnctt aatgggtggc tctgncnttt caannggaac cttngcnctn gntatgattg
                                                                          540
                                                                          600
ccacccegga naggggtggt ttggccttcc ntcttgtann aatctteneg gnttgttgga
                                                                          660
anggtnntct taggggatng ttccaatggg gaccgnaanc ttccagccna ggcaccaaan
                                                                          720
cnttggttta ncccccacnn aaaantanag gggncngqgt
                                                                          760
      <210> 157
      <211> 668
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(668)
      <223> n = A, T, C or G
```

```
<400> 157
ggtacccagt agtcattcag gaacaggttg ttcagtttcc atgtagttga gcggttttga
                                                                         60
grgagtttet taaacetgag ttgtegittg attgeactgt ggietgagag acagtttgit
                                                                         120
ataatttetg ttettttaea tttgetgagg agtgetttae ttecacetat gtggteaatt
                                                                         180
ttggaataag tgagatgtgg tgctaaaaag aatatatatt ctgttgattt gaggtggaga
                                                                        240
gttctgtaga tgtctattag gtctgcttgg tgcanagctg agtcaattcc tggatatcct
                                                                        300
tggtaacttt ctgcttgttg ntctgtctaa tattgacagt ggggcgttaa agtctcccat
attattgtgt gggagtctaa tetetttgta ggtetetaag gaettgettt ataaactggg
                                                                        360
                                                                        420
tgctcttgat tgggtgcaat atatttagga tagttagctc ttcttggtga atggancett
                                                                        480
taccaatatg aatggcetee tteettttga eettgtgggt taaagetggt tatngaaact
                                                                        540
ggatggance etgettttt tggtteattt ettgnagggt eeteageett attttanenn
                                                                        600
gnggetttgn ceencnteeg eggenttaag ggaaccaene tgngegteta ngancaetgg
                                                                        660
caactggg
                                                                        668
      <210> 158
      <211> 737
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (737)
      <223> n = A, T, C \text{ or } G
      <400> 158
tttttttaag ggtcaatgtt tacatttttt tcatataaat atcaagttgt cagcaccatc
                                                                         60
tgttgaaaaa aatctttgta atggctaatc ttttatgtca ttagatttga taatagttta
                                                                        120
agaatttttg ttcctatatt catgagggtt gctttccttt aacttttttg ttttgtaatg
tctgtgtcag gntttactat tagaacaata ctagtctagt aaaaaaaaa anaaacaaaa
                                                                        180
                                                                        240
aactancaag tgtntctccc cttctattta taanaanggn gttacttctt ccttaaatgg
                                                                        300
nnaaattatg agngaaactt ggagtatent tgenggantg gaagttteet tgtggaaaga
attttatnat nattacattt caatagtnee genteeetge negggeggnn nteaaaggeg
                                                                        360
                                                                        420
aatncagcaa attgntggcc gntactnngg accaacntcg gnccatnntg gggnancang
                                                                        480
tcaanctgtt ctngnnaatt gtncccttcc aatncccaca nanaaccgaa cctaaatgga
                                                                        540
accongggge tantaangne taccnntatt gngnggetnn geeettnint ggaaactgnt
                                                                        600
cnacenttat aatggeece enggaaggnt tntttggeet tetnntneaa anetggengg
                                                                        660
nttntgtgna ggttatctna ntggatgttc cacgggaacn gaanatntan ncagtggacn
                                                                        720
aaanntnntn ttttnct
                                                                        737
      <210> 159
      <211> 739
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (739)
      <223> n = A, T, C \text{ or } G
      <400> 159
cgagggtaca ctgtgagaga ataacatgga cttgatatgg catcacactt gttttaaagc
                                                                         60
aaaaaaaag aaaaaaagaa aaaaaagaaa gtacagttaa aaagtaagca ttgtagtaaa
```

```
tagtggattc tctggtgtgt attttttatc tcagtgttga aaattggaaa agaatgggct
                                                                        180
gaagtetaaa aactggaata atgaaggaca etaaatgeet ttattgtaga taetatgttt
                                                                        240
gtaagtctat agctaagcaa cttaagccaa aaaggtcttt caactgaagc tttaatcaac
                                                                        300
ttattttgga gatgttotot toottatoto atgogtoato cotaaaataa taagatacat
                                                                        360
gggatcaaat aaccettgce ttttcaacac aaatcagttg gaaaattatg ggttgagtce
                                                                        420
tgttgctgcc atggttctgt tctcaaaatg agtgtgtatg acatcccatc tatgtaatag
                                                                        480
gctacetttt tggctcttgg aactttgtcc tgccggccgg ccnttaaggc nantcnacca
                                                                        540
ctggcggccg tactatgggn tccagctcgt ccaaccttgc tatcntggct acttttctgg
                                                                       600
ngaatgtate cgtncatece caetteaneg gagetaangg aanentggge ctatgggget
                                                                       660
actecatatg ctngcenetg enttenangg aacnegente ttaanatgea eeenggaagg
                                                                       720
gtngtngcct tenttettt
                                                                        739
      <210> 160
      <211> 802
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(802)
      <223> n = A, T, C \text{ or } G
      <400> 160
cgaggtacag cagagacctt cctgcttttt actggggact ccagattttc cccaaacttg
                                                                        60
cttctgttga gatttttccc tcaccttgcc tctcaggcac aataaatata gttataccac
                                                                       120
taaaaaaaa aaaaaaaag tacgcggggg cccattgttt ttgtaatctc tgaggagaag
                                                                       180
cagcagcaaa catttgctag tcagacaagt gacagggaat ggattccaaa caccagtgtg
                                                                       240
taaagctaaa tgatggccac ttcatgcctg tattgggatt tggcacctat gcacctcag
                                                                       300
aggttccgag aagtaaagct ttggaggtca caaaattagc aatagaagct gggttccgcc
                                                                       360
atatagatte tgeteattta tneaatatga ggageaggtt gaetggeeat negaageaag
                                                                       420
aatgcagatg gcagtgtgaa gaaagaaaca tatttacctt taaagcttgg tcccttttna
                                                                       480
tegacenaag tggteegaca agettggaaa attactngan aaageteaat nggaetatgt
                                                                       540
gactettttt aataatttee anggntttaa accegtgagg actttteece egntaaatgg
                                                                       600
aaagtatttt genannggae ttgaetteee ggngeentaa gngaatteae caetgggggg
                                                                       660
gnttagggtc cnnntggnca anttggnaaa ngggtaatnn cntgnaatgt teeteateee
                                                                       720
aantngccgn ataantaacc gggcaaaggg cccaaatggn gccctccttn nngaatnanc
                                                                       780
cctntannna ancgggggg gg
                                                                       802
      <210> 161
      <211> 214
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(214)
      <223> n = A, T, C or G
      <400> 161
acttttnntt tattcnttat ttttgggacc tgctctcact gtccacccag actggagtgc
                                                                        60
antggcacca ttatagctna ctgcagcctt gacctnntgg gctcaagtga tcctnctgtc
                                                                       120
tacaccccc aagnatgntg tgacattatg cttggataat acttgtatnt tangtaaaga
                                                                       180
cagggtettt cenatnnace nggnagatet naaa
                                                                       214
```

```
<210> 162
      <211> 304
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (304)
      \langle 223 \rangle n = A,T,C or G
      <400> 162
acttaggaat acaactatat acatatgatt ttatttttaa gaccatatta tatttgggta
                                                                             60
totactaata tittgtataa agcaatittt tgttccatta cgtgactttt tgttttattg
                                                                            120
tatatgtaat ttaacacaca ataaagggta aagttgcttc cccaaaccac acttttaatc
                                                                            180
aaaacctaga atcatctgca gtccttgtta aaaatgcagg tttctagaac cctctgaagt
                                                                            240
tctgattaaa taaatttatt gcaaatcaaa naaaanaaaa aaaaaaaaaa agncccggg
                                                                            300
gnta
                                                                            304
      <210> 163
      <211> 461
      <212> DNA
      <213> Homo sapiens
      <400> 163
actagageca gteateetta acaaatettt teacatttta tttettteae atgtagteat
                                                                             60
cttcaaaaag gaaagatttg gaattttaga aaaggggcaa ctcttcttt tagcattctc
                                                                            120
atcagaaagt cacaaaaatc gatggaatca tttccactgg gaagattgac cttttgtatt
                                                                            180
tatttgtggg gtaaattaat aagcatteca gatgettgca getteetgea teeaggagat
                                                                            240
gctgtgttcc ccgtgatgca gctggaaccc aagctgcagc aggagatgca agtttcagga
                                                                            300
tgttccccac tgagctggag gaatatctac agcagtgatg cttgaaattt tgtatgaatt attttgtcgc ctaccctttt cctccaaaca aaaattagag gattatttaa tccttgggat
                                                                            360
                                                                            420
cttccccttt ttgagaaata aagtttttat caaaaaaaa a
                                                                            461
      <210> 164
      <211> 345
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(345)
<223> n = A,T,C or G
      <400> 164
tttttttgag acaaggtett actetgteae eeaggetgga gtgeagtgge atgatettgg
                                                                             60
ctcactgcac cctctgcatc ccaggitcaa gtgattctcc tgtctcagcc tcccttgtag
                                                                            120
ctgggattac agccacttgc cactgcaacc ggctaatttt tgtattctta gtagagatgg
                                                                            180
ggttttacca tgttggccag gctggtcttg aactcctgac ctcaagtgat ccacctgcct
                                                                            240
ccatgtccaa agtgctggga ttacaggcat gagccaccac ccctggccta agtcattaat
                                                                            300
ttaaaaaatg ttattaggat gancgacctg ccgggcggcc gntaa
                                                                            345
```

<210> 165

```
<211> 385
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(385)
      <223> n = A, T, C or G
      <400> 165
actgaaacag aaactntacc caattgcagt ccatatgttt tctgggatcc cggagttccc
                                                                        60
tttcaacaat gtaaaataca nacttaggtc aaaagttccc atgtctgaga aaactcaagc
                                                                       120
caaatcagtt ctcctccaaa gttgacagga tttatgcttt aaaaatagag atacagaatt
                                                                       180
ctctttggaa agatctacca aattcctgta agaaacagtc tacccaaagt aggggaaagg
                                                                       240
ctatatgana agttcaaggc acttcttaaa aatatatctt aggttttagg gaaaggaaac
                                                                       300
agacaagttt ccagacccgt gggtggaatg gatgtagcag atcactgaga ggttacaagc
                                                                       360
gccgacctng gccgngacac qctan
                                                                       385
      <210> 166
      <211> 745
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(745)
      <223> n = A,T,C or G
      <400> 166
tttttgacga tgtctctcaa caatacctga agtttctcat actcatcatc ccaagtctga
                                                                        60
aaaacttcaa agcatgctac cataactttt tcaaattctt cataagcaac atgcatcaat
                                                                       120
ttcctagtgc ccaatacttt gagtaattga gaactcaagt ctcttgaaat tgcctccacc
                                                                       180
aaacgcagtg ccctctgaat aggatatttt gtgtttcgga tctttctcaa atcccgcgta
                                                                       240
ctttgagaag ctgaggcggc agatcacttg aggccaggag ttcgagacca gtctcgtcaa
                                                                       300
catggcgaaa ccctgctcta caaaaaaaaa aaaanaanaa aaattagcca gacatggngg
                                                                       360
cccacatetg tagteccage taettganan getgaggeat gagaataget tgacetggaa
                                                                       420
nggcaaaggt ttantgancc caaactgngc ctggattcca atnnggngga cccagtgana
                                                                       480
tttgtctcaa aaaaangaaa ggaaaaaaga gcccgncgga aggaaggatg gattgangga
                                                                       540
aaattgtggc ctccnnnnaa aggnccaang gccctnangt ttctttgaat agtttccctn
                                                                       600
gcenttetta ngggeetnng cettttten nnetggegaa eetaggnatt cacatgggg
                                                                       660
ttangacnec geenetggga naggaaagtn etggaagnne nenteecaat anegnntang
                                                                       720
aacgggcngn ggannaattt tttnc
                                                                       745
      <210> 167
      <211> 623
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(623)
      <223> n = A,T,C or G
```

```
<400> 167
accagecact gcaaaacat gccaaattgt aaagaccate gaggetggga agaaactgca
                                                                              60
tcaactaacg agcaaaataa ccagctaaca tcataatgac aggatcaaat tcacacgtaa
                                                                             120
cactattaac ctgaaatgta aatggactaa attctccaat taaaagacac agactggcaa
                                                                             180
attggataaa gagtcaagac ccatcagtgt gctgtattca ggagacccat ctcatgtgca
                                                                             240
gagacataca taggctcaaa ataaaggaat ggaggaagat ctaccaagca aatggaaaac
aaaaaaaggc aagggttgca atcctagtct ctgataaaac agattttaaa ccacaaagat
                                                                             300
                                                                             360
caaaagagac aaagaaggcc attacataat ggtaaaggga tcaattcaca agaagggcta
                                                                             420
ctattctaaa tatatatgca cccaatacag gacccccaga ttcatgaagc aaatccttga
                                                                             480
gattnecaaa ggattaacte enecengtat tatggagaet tneacceact ntnacettte
                                                                             540
cegatettgn cecaaagtae enggttteee gaattgaetn gtttgneann gggetattaa
                                                                             600
tttngaattt cncccaaaaa aaa
                                                                             623
       <210> 168
       <211> 703
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (703)
       <223> n = A, T, C \text{ or } G
       <400> 168
ggtactccct gtttgctgca gaatgtcaga tattttggat gttgcataag agtcctattt
                                                                              60
gccccagtta attcaacttt tgtctgcctg ttttgtggac tggctggctc tgttagaact
                                                                             120
ctgtccaaaa agtgcatgga atataacttg taaagcttcc cacaattgac aatatatatg
                                                                             180
catgtgttta aaccaaatcc agaaagctta aacaatagag ctgcataata gtatttatta
240
                                                                             300
gcaaattata tttttgctgc tgatatatta gaataatttt taaatgtcat cttgaaatag
                                                                             360
aaatatgtat tttaagcact cacgcaaagg taaatgagca cgttttaaat gtgtgtgtgc
taatttttte cataagaatt gtaaacattg actgaacaaa tacctatatg gattggtaat gacttatgag caanctgett ggecagacag ttacccaaac tttatatatn tnngaaggta tacactgnga aatetetgge taancgaatg entecagggg taanngggtn tggntggant
                                                                             420
                                                                             480
                                                                             540
                                                                             600
aaanaatgcc ctgcaaaaaa aaaaaaaaa aagccttccg nggccttnaa nggaatcnnc
                                                                             660
angggnntnn ggccactggc cactggnaaa ngnaacgtct gga
                                                                             703
       <210> 169
      <211> 609
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(609)
      <223> n = A,T,C or G
      <400> 169
acgtccatct tccagctgct tgccagcaaa gatcagtctc tgctgatcag gaggaattcc
tteettatee tggatettgg cetttaeatt ttetategta teegagggtt caacetegag
ggtgatggte ttaccagtca gggtettcac gaagatttge atcccacete tgagaeggag caccaggtge agggtggact etttetggat gttgtagtca gacagggtge gtecatette
                                                                            120
                                                                            180
                                                                            240
cagctgtttc ccagcaaaga tcaacctctg ctggtcagga gggatgcctt ccttgtcttg
                                                                            300
```

```
gatetttgee ttgacattet caatggtgte acteggetee acttegagag tgatggtett
                                                                       360
accaagtcag ggtcttcacg aagatctgca teccaectet aagacggage accaggtgca
                                                                       420
gggtggactc tttctggatg ttgtaatcag acanggtgcg ttcatctttc actgnttcca
                                                                       480
caaaaaacaa cetetgetgg canganggat cetteetine tiggactitg cetgacatte
                                                                       540
tnatggngta ctccgctccc ttcaaaggga tgncttacan tcanggnctt acnaaaattt
                                                                       600
cntccnctt
                                                                       609
      <210> 170
      <211> 617
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (617)
      <223> n = A,T,C or G
      <400> 170
acaaagaaca tgtagctata ggaaataata gtgtaaatag cagtatataa actggcccat
                                                                        60
gtaaaataca aaaatattca ctgaagtcag gttttctata aaacagtgtt tattagaggt
                                                                       120
attttactat gaatcaggca tataatctga atgtagaaac ttttagaaat attaacagca
                                                                       180
ttcagtcagt gccatgcact tgtgcttcca attattttt taaagctgct ttgttttgac
                                                                       240
tcatgtgaaa tagttaaggc ctacattctt atacacatta tccatcttac aaggttaaca
                                                                       300
attttacact aaaacacagt ttaaattaaa aacgattttg aaaaattaca tctatattta
                                                                       360
atccctaaga agtgttttaa gctggtaatg cagctcgctg tagctctaag agaggggtta
                                                                       420
gtcaggaatc tgatcttgag ccataaangg tttcaggcta aacaaagaac aaatttaagt
                                                                       480
gacagaaaat attataattn caatatactc agttttttgg tataaaatac cctgctagca
                                                                       540
tgccactggc tatattgngg gcataatata aaatgncggg gggggggatg gancctccaa
                                                                       600
gncaaanttt ggaccca
                                                                       617
      <210> 171
      <211> 621
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(621)
      <223> n = A, T, C or G
      <400> 171
acagtatggg ggttgtaaat tggcatggaa atttaaagca ggttcttgtt ggtgcacagc
                                                                        60
acaaattagt tatatatggg gatggtagtt ttttcatctt cagttgtctc tgatgcagct
                                                                       120
tatacgaaat aattgttgtt ctgttaactg aataccactc tgtaattgca aaaaaaaaa
                                                                       180
aagttgcage tgttttgttg acattetgaa tgettetaag taaatacaat tttttttatt
                                                                       240
agtattgttg teetttteat aggtetgaaa titttettet tgaggggaag etagtetttt
                                                                       300
gcttttgccc attttgaatc acatgaatta ttacagtgtt tatcctttca tatagttagc
                                                                       360
taataaaaag cttttgtcta cacaccctgc atatcataat gggggtaaag ttaagttgag
                                                                       420
atagttttca tccataactg aacatccaaa atcttgatca gttaaaaaat ttcacataac
                                                                       480
ccacttacat ttaccaactg gaagaataat caatctctca agcatgggat tattagaatc
                                                                       540
aacantttga aagctgtcct tgaaggctaa taaaaaagnt tgtctaacct ttcatgaggn
                                                                       600
cttnttntta ctnccttacn g
                                                                       621
```

```
<210> 172
      <211> 399
      <212> DNA
      <213> Homo sapiens
      <400> 172
actcaaaatt acacatttgt ttaaataaat atccacacaa attctcagtt acatcaagta
                                                                            60
gctggtttat atttagatta tctcaagtag gggggaataa ccatgtgtag gaattcatag
                                                                           120
aaaaataaac aatcagctga agaggtctaa gaaaatgctg acttttaaaaa tttcacttat
                                                                           180
tttccttgaa gttttctacc cttcccatcg atgataaacc aagatcatgt aatggaaaat
                                                                           240
ttcaaaccag ggctaaattc taaaqtaaag cttcaattca agcccttccc ccaaqaqaat
                                                                           300
taattttcct gatttctctt tctctcacat ctaaqqaqaa cattttaqqc aqttaaattt
                                                                           360
cagaacttca aggtttcatc agggtcacct ttatgtacc
                                                                           399
      <210> 173
      <211> 616
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(616)
<223> n = A,T,C or G
      <400> 173
actttgtgga taagaaaatg gaggaacaca tctgatggag agtgggcatt tgacaacaat
                                                                            60
ggaacaggta acctgcatgt aaaatcaaaa tataagtgtc tttttaagag ctgaaagctg
                                                                           120
ctgctggtca ttcattaatg tgtcagacat ttaatcagga tgctggacct tcaaaataac
                                                                           180
tgaaaaaaga accaagaaaa ggcgtttttg ttttcaacaa actttactaa ataaccctgg aaaggcaatg aacgatctga caatttaagc tctaatgatt taaagctcag ctagaagaaa
                                                                           240
                                                                           300
gtgaggcatg acatatactg tcaacggagg gtgaaggagg canatttctg gaaatgcaat
                                                                           360
gatcccacca tttgcttcaa ngagaaacct gcanacatat tttcangtct tgntaagtna
                                                                           420
caactginta titigiaatca atcattingg aaaagtcigc tatgiaactt angnoactgi
                                                                           480
gccccnacc accgatgaaa aggaaaaacc cctgacacca ggaaaatcct tccatcctca
                                                                           540
aanaaattaa gngaccaacn tttaaagaaa aaaaatnanc ccncctctnt ttacaaatnt
                                                                           600
ttcntccaaa tnttcn
                                                                           616
      <210> 174
      <211> 631
      <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
       <222> (1)...(631)
      \langle 223 \rangle n = A,T,C or G
       <400> 174
ggtacgcggg gacacgcacg ccgggcgtgc cagtttataa agggagagag caagcagcga
                                                                             60
gtettgaage tetgtttggt gettiggate calttecate ggteettaca geegetegte
                                                                            120
agactccagc agccaagatg gtgaagcaga tcgagagcaa gactgctttt caggaagcct
                                                                            180
tggacgetge aggtgataaa ettgtagtag ttgaettete agecaegtgg tgtgggeett
                                                                            240
gcaaaatgat caagcettte ttteatteee tetetgaaaa gtatteeaae gtgatattee
                                                                            300
```

. . ... .....

```
ttgaagtaga tgtggatgac tgtcaggatg ttgcttcaaa agtgtgaagt caaatgcatg
                                                                              360
ccaacattcc agttttttaa gaaagggaca aaaggtgggt gaattttctg gagccaataa
                                                                              420
ggaaaagett gaagecacca ttaatgaatt aatetaatea tgttttetga aaacataace
                                                                              480
accattggct atttaaaact tgtaattttt ttaattttcc aaaatttaaa tttgaanact
                                                                              540
taaccccant tgccatntgn gtgacaataa aacattatgc tacccntttt aaaaaaaaaa
                                                                              600
aaaaaaaaa agteetgeee ggeggeeete a
                                                                              631
       <210> 175
       <211> 261
       <212> DNA
       <213> Homo sapiens
       <400> 175
acgaacctac agttttaact gtggatattg ttacgtagcc taaggctcct gttttgcaca gccaaattta aaactgttgg aatggatttt tctttaactg ccgtaattta actttctggg
                                                                              60
                                                                            . 120
ttgcctttgt ttttggcgtg gctgacttac atcatgtgtt ggggaagggc ctgcccagtt gcactcaggt gacatcctcc agatagtgta gctgaggagg cacctacact cacctgcact
                                                                             180
                                                                             240
aacagagtgg ccgtcctaac c
       <210> 176
       <211> 616
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (616)
      <223> n = A,T,C or G
      <400> 176
cgaggtactc tgccttttag gagatgaggt aagacatata catagatggc ttttactagc
                                                                              60
caaggcaatg taaatggact aagattetea tgtgacttga ggttatetga tgaatttatt
                                                                             120
ctcttcaaaa ccacctacct ttagagggca tgtttaaccc ctctctttat ttaaggaggg
                                                                             180
agagaaaaac acatgtaacc agaattcaga gtgggttact caacctaaga gaacatacgg
                                                                             240
agttetettt gggaaaacaa caagactaca gtgtteaett egeaceatga agtggeaete
                                                                             300
ctgttatggc tgtcagagtc ctctcacttc ttatgaaagg atgcatctga ttctgaaatt
                                                                             360
actgatatat tegateagtt anggatgttt taaaaagtga aaacaaatge cacacataca
                                                                             420
ctttctagct ttcttgaaat caccegacac attccaaaaa tagagaattc cctattactt
                                                                             480
ttagagaaat ttccatatan tcttggtnaa gaanccagtt gngcntattc caatttcagg
                                                                             540
gtcttggttt ttgcccaaac ccaagigttt contntttta nggcttttca tggccgattt
                                                                             600
naaaccttnt ttgtgg
                                                                             616
      <210> 177
      <211> 632
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (632)
      <223> n = A,T,C or G
      <400> 177
```

```
cgaggtacag gtcagagtct tcttttcttt tctttttgag atggagtctt gctctgttgc
                                                                           60
cagactggag tgcagtggtg cgatctgggc tcactgcaat ctccacctcc cgggttcaag
                                                                          120
cgattetect geeteageet ecegagtaae tgggaetaea ggtgtgegee accaageeea
                                                                          180
gctcattttt gtatttttag tanagatggg gtttcacggt gttggctagg atggtctcga
                                                                          240
tctctggtca gaagtctttt ctgtaaatat ccttggtaaa gaagcaattt tagactgtag
                                                                          300
ctgttgcaaa tgctttaagg aagaagcaaa acaactgtca gtcttcctga aatgaaaaaa
                                                                          360
ctacaccagg gctgctatat caaagcaacc ccaaccagca cttcaatcat gatgcccaca
                                                                          420
gtggccccac tgagaaacca agaaaagttn cagatacaaa actgngatgc tcttgctatg
                                                                          480
gnaatattgc nggcngtanc caagttagaa accaaacaag cntanggccc cgttnttttt
                                                                          540
tggcgtgatt ttggcaanaa aaaaactgg gngngtggtg ngggttccca ttgtacccc
                                                                          600
aaaaaacttn gggatgggtt aaagcccnng gc
                                                                          632
       <210> 178
       <211> 611
       <212> DNA
      <213> Homo sapiens
       <220>
       <221> misc feature
      <222> (1)...(611)
      <223> n = A, T, C or G
      <400> 178
actitntttt ttttttttt ttttttttg ggatttagtt tttatttcat aatcataaac
                                                                           60
ttaactctgc aatccagcta ggcatgggag ggaacaagga aaacatggaa cccaaaggga
                                                                          120
actgcagcga gagcacaaag attctaggat actgcgagca aatggggtgg aggggtgctc
                                                                          180
tcctgagcta canaaggaat gatctggtgg ttaagataaa aaacaagtca aacttattcg agttgtccac agtcagcaat ggtgatcttc ttgctggtct tgccattcct ggacccaaag
                                                                          240
                                                                          300
egetecatgg cetecacaat atteatgeet tettteactt tgecaaacae cacatgettg
                                                                          360
ccatccaacc actcaatctt ggcagtgcag atgaaaaact gggaaccatt tgtgttgggt
                                                                          420
ccaacatttg ccatggacaa aatccangac ccgtatgctt taagatgaaa ttctcatttc
                                                                          480
aaatttette ecataaatgg aettgeenea tgeeatnttg ggtgtgaagt neeneettge
                                                                          540
ncataaccct ggaatatttt tgaaacagaa ccttttacca atcnttttt catgttaaaa
                                                                          600
acnaaaattt t
                                                                          611
      <210> 179
      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (611)
      \langle 223 \rangle n = A,T,C or G
      <400> 179
acctcaattt tatcatttta gagtatttgt tagaatagga tctctccaaa atcaaacagg
                                                                           60
atcaatctgg tcacgtctaa tcctaagaca aaacactatg taaaattttc ctgtatctaa
                                                                          120
atgttgccct ctaggtaaat ctgtgatatt ttagagactt tcttttgtgg aaaaggtaat
                                                                          180
ctgataaatg ggaagagatc atcagacaag ttcacaaata accattattt ctgcagaatt
                                                                          240
cagttgaagt tggttttttg taaatgetta ttgggaattt ctaaageact gaettggaga
                                                                          300
ggccaagage etecateaat ecetgettgg atagecaete eegttaetae tgetaggtea
                                                                          360
gggtetacag atgtgttggg atetttteca aagaaetett gaatgaettg aeggateega
                                                                          420
```

.....

```
ggaataccaa tggagccccc aactaaaacc acctcatcaa tctcagtctt ttncaqqtqq
                                                                              480
nettetteaa teteetgaat gggaeetegg eegeaneaen etanggegaa tteeacaeet
                                                                              540
ggcggccgta ctaatggatc caactcgnac caacttgggg aacatggcta gtnttcnngg
                                                                             600
ggaaatgttt c
                                                                             611
       <210> 180
       <211> 621
       <212> DNA
       <213> Homo sapiens
       <221> misc feature
       <222> (1)...(621)
       <223> n = A, T, C \text{ or } G
       <400> 180
accettaaac tggcaggaca tttttgaaat cacaaatttg cacataaaga atgtcacgaa
                                                                              60
cagccatgta tccatataca gcaatcaaat aaggaactta tgacctaaag caaaggtaaa
                                                                             120
ctttcttgaa acttaacatt ctataccaac taggcaacct ctgcccagga tgagagttgg
                                                                             180
atttttcaaa aacctctaat ttaatagtgc agcatttcgt tttccctgat ggcctgtgtt
                                                                             240
tcacagcagt ttttaaaaac tgcttgttca actatagctg cagcctatat cccagctatg
                                                                             300
gaaaaaaaag taaatettag ticaattttt gecagtigti teigtattta aattiaaaaa aaaacacact teegetggge aggittagag ggitattate aagietgige ataactaaaa
                                                                             360
                                                                             420
gttcaaagca aattcaattt tgcttaangg aacattgnna aagnacaatt cttggnanta
                                                                             480
catgcctcgt tgatccattt naancatana aaattcaccc ttgtgtactg gttcaagaaa
                                                                             540
aaaaccgatt tgacagttaa acatnttaaa anccccaacc tntgaagttc aaccaaactg
                                                                             600
ganttttgtt cctcgcccga c
                                                                             621
      <210> 181
      <211> 606
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(606)
      <223> n = A,T,C or G
cgaggtacag accagagaca aagcaagaga agaagcagag actgttggcc cgggccgaga
                                                                              60
agaaggetge tggcaaaggg gacgteecaa cgaanagace acctgteett cgageaggag
                                                                             120
ttaacaccgt caccaccttg gtggagaaca agaaagctca nctggtggtg attgcacacg
                                                                             180
acgtggatec categagetg gttgtettet tgeetgeeet gtgtegtaaa atgggggeee
                                                                             240
cttactgcat tatcaangga aaggcaagac tgggacgtct agtccacaag gaagacctgc
                                                                             300
accactgtcg cetteacaea ggtgaacteg gaagacaaag gegetttgge taaaetggtg gaagetatea ggaccaatta caatgacnga taenatgaga teeeceteet ggggtggeaa
                                                                             360
                                                                             420
tgtcctgggt ctaaatctgt ggcttgtatn gccaacttcn aaangcaaag cttaaaaact
                                                                             480
tgcncttaac tngggtnaat gtactncccg gcggccgttg aanggcaatt caacacattg
                                                                             540
cggccgtcta atggntcanc ttggnccaac ttgggnaana tggnaaannn ttcttgggna
                                                                             600
atttnn
                                                                             606
      <210> 182
      <211> 610
```

```
<212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (610)
      <223> n = A, T, C \text{ or } G
      <400> 182
ggtactcata aaaaaagtct taccccaaaa ttgcaaacaa atacattaaa agattagaag
                                                                      60
120
aaggtcccca ttcagcaaat actttgtaaa gtatggcctg tatgtaaata gtgctaaatc
                                                                      180
aaggactttt tagcagaaaa ttgctcggtt cttttatcta aggcttgaat ttgtaaagtg
                                                                      240
aaggcataaa agttaccaaa cattaagtaa ctcttaaaat ggcacacagg ttttaaagct
                                                                      300
attggttttt ccttcctaac tctctgaatt tttcccatgg cctttgtaga tcaactattt
                                                                      360
caaacgtatt ttacaccagc aactctcaac atacttgtct ttcagatatg tcatcagtca
                                                                      420
tgtctaacag gccaatagcc aaataacnga tttaaaacaa tncttaacta gctagcagga
                                                                      480
cattactttg gatctgctta ctgcaactga ctatttgtaa gcttaaaatc antttaatcc
                                                                      540
tgatacagaa acctcatctg cncatacntt actttggcct tcaaccttta aaaatactta
                                                                      600
atcccccqnc
                                                                      610
      <210> 183
      <211> 608
      <212> DNA
      <213> Homo sapiens
      <221> misc feature
      <222> (1) ... (608)
      \langle 223 \rangle n = A,T,C or G
      <400> 183
60
tttttttggg agncagctnt ttaattaggn tcttaaaaca tttaaaacnc caatttgnga
                                                                      120
ggataaattc cattcgtcan ancaaacnca aatcgcaggt anccctggan ctgaggaata
                                                                      180
netttgattt ttggnaaaat ttgngagtee acagettint gateaaintt genetgetee
                                                                      240
gnaatctcat atttctnttt ttctgngncg aaaatctcac cttcctggng tntgggcttc
                                                                      300
cgcagcttnt tntttttgaa gtaagcatca ataaaangtt ttgggatttt tacattgctg
                                                                      360
aaatccattt tgggtgaagg ggcaatgaca aatttntngn gtnttctttt taaaagaacc tcattggggg ccnaaggncc cncccaaatt ataaaccct ttccccctgg tttangnaaa
                                                                      420
                                                                      480
ccccctttg ccctgngggg nccangagga taaanaaagg ccccggggaa gctggccca
                                                                      540
ntttttcccg ccgncgaagg gttttgccgg ctaaaanttt tngggcattt nnngggnaat
                                                                      600
tttggctt
                                                                      608
      <210> 184
      <211> 622
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (622)
      <223> n = A,T,C or G
```

```
<400> 184
acagecetga tgcaaagttt cagageatga ceageaagtg geeagetgtg tgggteaaga
                                                                           60
teageteeag etgggtetge eteetgettt acgtetggae cettgtgget ceaettgtee
                                                                          120
tcaccagtcg ggacttcagc tgaacctctg agtgccaagg acaccactgg aactcacaaa ggtctccttc accgaaaacc catatacctt ttaagtttgt ttcaactaaa atattaagtg
                                                                          180
                                                                          240
aatgetttge aagittgaet gtatgeaggt ttatateaag aaggtgagat tgaataatge
                                                                          300
ttgatgcaga atcgaaactt ctcatttatc tgnatattat gtttacttct aaggatatag
                                                                          360
cacaaaggga acattttttg tttaaagtga actacagetg tgctgtgaag agagttcttt
                                                                          420
ataaagcctg taggtctttt aactttggtt aaaatgtaag ataggaaaat gttggatatt
                                                                          480
tgaggcntgc ctaatatatt tatattggag natcctttna aagccaaaaa aaaaaaaaaa
                                                                          540
aaaaaaaagt nccttggccg gaccncccta aggggaattc cacncactgg gggccgtntt
                                                                          600
atggatccaa ctcgnaccaa ct
                                                                          622
      <210> 185
      <211> 614
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (614)
      <223> n = A,T,C \text{ or } G
      <400> 185
acgeggggac agteceacee teacaegatt etttacettt caetteatet tgecetteat
                                                                           60
tattgcagcc ctagcagcac tccacctcct attcttgcac gaaacgggat caaacaaccc
                                                                          120
cctaggaatc acctcccatt ccgataaaat caccttccac ccttactaca caatcaaaqa
                                                                          180
egecetegge ttacttetet teettetete ettaatgaca ttaacactat teteaceaga
                                                                          240
cetectagge gacceagaca attatacect agecaacece ttaaacacec etecceacat
                                                                          300
caagecegaa tgatatttee tattegeeta cacaattett egateegtee taacaaacta
                                                                          360
agaggegice tigecetatt actatecate eteatectag caataatece atcetteata
                                                                          420
tatcccaaca acaaagcata atatttcgnc cactaagcca atactttatt gattctagcc
                                                                          480
ggagacetet nantntaace tggateggag gaaaceagta getaceettt accaatantg
                                                                          540
ganaagaaga tegnacettg gegggacaee ttangggaat teaaceaetg gnggeggtat
                                                                          600
atgggacccn ccng
                                                                          614
      <210> 186
      <211> 627
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (627)
      \langle 223 \rangle n = A,T,C or G
      <400> 186
ggtactgatt ttaaaaacta ataacttaaa actgccacac gcaaaaaaga aaaccaaagt
                                                                           60
ggtccacaaa acattctcct ttccttctga aggttttacg atgcattgtt atcattaacc
                                                                          120
agtettttae tactaaaett aaatggeeaa ttgaaacaaa cagttetgag accgttette
                                                                          180
caccactgat taagagtggg gtggcaggta ttagggataa cattcattta gccttctgag
                                                                          240
etttetggge agaettggtg acettgeeag etceageage ettettgtee aetgetttga
                                                                          300
```

```
360
tgacacccac cgcaactqtc tqtctcatat cacqaacagc aaagcgaccc aaaggnggat
agtetgagaa getetnaaca cacatggget tgecaggaac catatnaaca atggcagcat
                                                                           420
caccagactt naagaattta agggcatctt ccacttttta ccaaaacngn gaacaatctt
                                                                           480
tttenttact taacnaacht getteeatgg gageegggng naateeaate aagggeataa ceegggeett atttggenng atgggteang gnaataneet gaeeaggaaa eeeetgntte
                                                                           540
                                                                           600
cttgggggga antttgttgn nccccac
                                                                           627
      <210> 187
      <211> 256
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(256)
      <223> n = A, T, C or G
      <400> 187
                                                                            60
ggaccttttt ttttttttt ttttttttt ggaaaagaaa ggccttacat atttattact
gaatccagcc aaccaacgtg ttcataacag attcagagag gaaaacacgt cgaaatctcc
                                                                           120
                                                                           180
anatagtggt gacattttca gettgatatg gtaacatgat egtgacette anacagcata
aatatgtgtg ccatctcatg tgcaattcct tatanaccca gcttggttct tctccaatgt
                                                                           240
                                                                           256
ctccttttgg agttgt
      <210> 188
      <211> 523
      <212> DNA
       <213> Homo sapiens
      <220>
       <221> misc_feature
       <222> (1) ... (523)
       <223> n = A, T, C \text{ or } G
       <400> 188
                                                                            60
ggtaccacct acacccaaca agtcaatgag ggacttcttt ttaatttggt aggattttga
ctggttttgc aacaataggt ctattattag agtcacctat gacaaaaaat aggggttacc
                                                                           120
tagataatgc caaagtcagc atttgtcctg ggttcccttg tgtgatctgt ttggactatg
                                                                           180
                                                                           240
ttitetttie treteceaet tgeteageag ettgggette cattetaget ettttaceaa
                                                                           300
gattttttqtq tqaccatqtt qacttcattt ggattgccct ctttcaattt ccttgtgaaa
                                                                           360
acaccettaa etttetettt accettaget gaaatgitta cataactiet ggigatatet
                                                                           420
tttcatgatt ttatatctct taaaatggtg atggatgtga cacctcataa aagtgagctt
tgaactgtag ataactctta aagaaaatgt cattttanac aattaaaata tttgtgctca
                                                                           480
                                                                           523
aaaaaaaaa aaaaaaaaa gtcctgcccg gcggccgtcn aan
       <210> 189
       <211> 622
       <212> DNA
       <213> Homo sapiens
       <221> misc feature
       <222> (1)...(622)
```

```
<223> n = A, T, C \text{ or } G
      <400> 189
acaatttaat ttttctgctt gcccaagaaa caaagcttct gtggaaccat ggaagaagat
                                                                           60
                                                                          120
gaaaatgaga ctggcaaaga acaaatgctg aatctgaaga agaggacaac tttgggcaaa
taatctgcat acttttaatt gggaataaga tggaaaatat gaatgctaaa tcaaattttt
                                                                          180
taaaaaatac accacacgat acaactcaat acaggagtat ttcttctcaa attcttctag
                                                                          240
caccatcaac attetteaag tatetqaaat actattaatt aagcacettt gtattatgaa
                                                                          300
caaaacaaaa caaggacctc agttcatctc tgtctaggtc agcacctaac aatgtggatc
                                                                          360
acactcatgg gaaagtgttt tgaggtagtt taaacctttt ggaaggttgg gttttaaact
                                                                          420
tecetetgtg gaagatatea aaageeecaa gtggtgeeaa atggttatgg ttttattttt
                                                                          480
caattttaat ttgggtttct tccaaaggtg acatttccat acaaggggaa gggggtggaa
                                                                          540
aaaaaatcaa attttggggg accagggagg ataatnaact gtttgcaatg cttgacaacc
                                                                          600
ttttttttt gnccaantaa ca
                                                                          622
      <210> 190
      <211> 628
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (628)
      <223> n = A, T, C or G
      <400> 190
accactaata gggtgtatct cagaaactga attgaaataa gggaaaatag gattttctgt
                                                                           60
cetggttttt gaagattgtt ettgatteee ttgatteeea ggagagatte tetgacatte
                                                                          120
acgigicage cacitiggea eggaageett acagigiggg gaaccaaaac tiegigtete
                                                                          180
ctettteece gatgecatea geatagaett gaetteetta aacegagagt tttgatgtgg
                                                                          240
cettggcaac cetaaaatca getgtgttag gtaacaaaac teaggettte tgttgatgae
                                                                          300
atcgagatgg tgtcacttaa aagagccaag attcctgttt tcagtttgtg gattcatcct
                                                                          360
gctggtttta ctttagtccc tccatgtcaa agtgggcctg agaaaagctc atacatgcct
                                                                          420
catgtgaagt gtccacccc tctgaaaatc tttcttgttc aaaacancna cgacatatct
                                                                          480
tggtaacttt tacggtgact tttggangag gggagtttgg aaattgtaaa atgttatana tggtgcctat ttcctgctga angaaatgtt ttaaaaaagnn tntntaancn taatcnaatg
                                                                          540
                                                                          600
gttgggggn qaccttctac cnaanntn
                                                                          628
      <210> 191
      <211> 474
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (474)
      <223> n = A,T,C or G
      <400> 191
ggtacagccc tcaatctgtt cttcaagctc aagaacttca agacagctgc cacctttgct
                                                                           60
eggegeetae tagaaetegg geecaageet gaggtggeec aacagaeeeg aaaaateetg
                                                                          120
totgootgtg agaagaatco cacagatgco taccagotca attatgacat gcacaaccoc
                                                                          180
tttgacattt gtgctgcatc atatcggccc atctaccgtg gaaagccagt agaaaagtgt
                                                                          240
```

```
300
ccactcaqtq qqqcctqcta ttcccctqaq ttcaaaggtc aaatctgcag ggtcaccaca
                                                                       360
gtgacagaga ttggcaaaga tgtgattggt ttaaggatca agtcctctgc agtttcgcta
aagccccctt tgtgtgcatg ggtcaagtca ccatatgttc cccccaaaaa atgtgtctat
                                                                       420
                                                                       474
atotecttot aacaacacot toccotqoac tactottoaa atotngotot ntgt
      <210> 192
      <211> 234
      <212> DNA
      <213> Homo sapiens
      <400> 192
acgeggggt tggtgagtgg getectaceg accgaggttt aggeagegeg gggagetttg
                                                                        60
cgggttgcca tttgtaactc cggatectaa aattectgte etgttetetg tetettetag
                                                                       120
gttgggggcc gtcccgctcc taaggcagga agatggtggc cgcaaagaag acgaaaaagt
                                                                       180
cgctgqaqtc qatcaactct aggctccaac tcgttatqaa aaqtggqaaq tacc
                                                                       234
      <210> 193
      <211> 367
      <212> DNA
      <213> Homo sapiens
      <400> 193
ggtaccaata ccaccaattt tgtagacatc ctggagaggc aggcgcaagg gcttgtcagt
                                                                        60
tggacgagtt ggtggtagga tgcagtccag agcctcaagc agcgtggttc cactggcatt
                                                                        120
                                                                        180
gccatcctta cgggtgactt tccatccctt gaaccaaggc atgttagcac ttggctccag
catgttgtca ccattccaac cagaaattgg cacaaatgct actgtgtcgg ggttgtagcc
                                                                        240
aattitetta atgtaagtge tgactteett aacaattiee teatatetet tetggetgta
                                                                        300
gggtggctca gtggaatcca ttttgttaac accgacaatt agttgtttca cacccagtgt
                                                                        360
cccgcgt
                                                                        367
      <210> 194
      <211> 613
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (613)
      <223> n = A,T,C or G
      <400> 194
ggtactcttg gtttqtcaat qqqactttcc aqcaatccac ccaaqaqctc tttatcccca
                                                                         60
                                                                        120
acatcactgt gaataatagt ggatcctata cgtgccaagc ccataactca gacactggcc
tcaataggac cacagtcacg acgatcacag tctatgcaga gccacccaaa cccttcatca
                                                                        180
ccagcaacaa ctccaacccc gtggaggatg aggatgctgt agccttaacc tgtgaacctg
                                                                        240
agattcagaa cacaacctac ctgtggtggg taaataatca gagcctccgg tcagtcccag
                                                                        300
gctgcagctg tccaatgaca acaggacct cactetacte antqtcacaa ggaatgatgt
                                                                        360
aggaccctat gagtgtggaa tccanaacga attaagtgtt gccacagcga cccagtcatt
                                                                        420
ctgaatgtcc tctatgncca gacgaacccc catttcccct cataccctan taccgtcaag
                                                                        480
ggtgaacctt agctttctgc atgcagcttt aaccactgcc agtttcttgn tgatgatgga
                                                                        540
catcacaca cacaagactn ttatticaca tactgagaan aaagcgactt ntactgcagg
                                                                        600
cataactanc ngg
                                                                        613
```

```
<210> 195
      <211> 613
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (613)
      \langle 223 \rangle n = A,T,C or G
      <400> 195
acgogggcgc cagagtccct gaactotegc tttcttttta atcccctqca tcqqatcacc
                                                                            60
ggcgtgccc accatgtcag acgcaqccgt agacaccagc tccgaaatca ccaccaagga
                                                                            120
cttaaaggag aagaaggtga tggtgaggaa gaggatggag atgaagatga ggaagctgag
                                                                            180
tcagctacgg gcaagcgggc agctgaagat gatgaggatg acgatgtcga taccaagaag cagaagaccg acgaggatga ctagacagca aaaaaggaaa agttaaacta aaaaaaaaa
                                                                            240
                                                                            300
aggoogoogt gacctattca coettcactt toogtotnaa aatotaaacg togtcacott
                                                                           360
caataaaaag geceeegee eengggeag tgeeeecea aaataaaege gettteacea
                                                                           420
ccaaccaaac atgaaaattt tccacaaggg anggaaaaaa aaccaaacnt ccaaggcctn
                                                                           480
ttttttttta aaatactngg ccgcgaccac cctanggcga attccanacc tggcggccgt
                                                                           540
nttatggatc cnactcggac caacttgggn aatatggcat antggttctt ggngaaatgt
                                                                           600
atccctccat tcn
                                                                           613
      <210> 196
      <211> 296
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(296)
      \langle 223 \rangle n = A,T,C or G
      <400> 196
geggnggenn ggeegaegnn eteateaatg ttgtteggte ageeetteee taattacace
                                                                            60
tatconotac acatacatgo acatagacac acnontgaac noactgaana tatttootto
                                                                           120
aggtgtgtgt aaaatatgct gcttggattg aaattcannt gggattgatt agncaagtan
                                                                           180
cttganacct cacagtaatc ttcacacttn nccttacaca cctatgcagg catgttggga
                                                                           240
grangttaca atgttacttc agreeacagt ttatttctat acttqaqttc ttaaqt
                                                                           296
      <210> 197
      <211> 222
      <212> DNA
      <213> Homo sapiens
      <400> 197
acatggagga gaatgaccag ctcaagaagg gagctgctgt tgacggaggc aagttggatg
                                                                            60
tegggaatge tgaggtgaag ttggaggaag agaacaggag cetgaagget gaeetgeaga
                                                                           120
agctaaagga cgagctggcc agcactaagc aaaaactaga gaaagctgaa aaccaggttc
                                                                           180
tggccatgcg gaagcagtct gagggcctca ccaaggagta cc
                                                                           222
      <210> 198
```

<211> 539

```
<212> DNA
      <213> Homo sapiens
      <400> 198
cgaggtacta catatttcag cactaaggcg gttgcttcac tttatatcta tataaaaaaa
                                                                        60
gtggtaaaaa tetttteett ttgtgeagtt gaacceatee tacatteaga tteteteaag
                                                                       120
cactaataaa atacttattt ggttgaggaa gatttaaggc aagttcgggc ccttccaaag
                                                                       180
geactgtgag actececce cactececgt tattgetaca tgtetttata etegagtatg
                                                                       240
tcacagtaga actggtggaa taagcaaaca cttttttgct agtttataaa gttggaatta
                                                                       300
gaaaagcatg ccacatttca gcctgattgc aaagtatgtg gtcatttttt tctttgaagt
                                                                       360
tggatgggct acaaccttta tacattctaa gaaaactcat aggatgttcc tcaaactact
                                                                       420
tccacagcat caagatcgat ttctgtcaag aaatcatgca atctttcaaa atttacgtaa
                                                                       480
acaaggaaag aaattaatga aataaatatt acatacaatc tottaaatta agaatttgt
                                                                       539
      <210> 199
      <211> 626
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(626)
      <223> n = A, T, C or G
      <400> 199
cgaggtacaa gatgtccaaa tattgcgaag atctatttgg ggatctcctg ttgaaacaag
cacttgaatc acatccactt gaaccaggca gggctttgcc atcccccaat gacctcaaaa
                                                                       120
gaaaaatact cataaaaaac aagcggctga aacctgaagt tgaaaaaaaa cagctggaag
                                                                       180
etttgagaag catgatggaa getggagaat etgeeteece agcaaacate ttagaggacg
                                                                       240
ataatgaaga ggagatcgaa agtgctgacc aagaggagga agctcacccc gaattcaaat
                                                                       300
ttggaaatga actttctgct gatgacttgg gtcacaagga agctgttgca aatagcgtca
                                                                       360
agaaggette agatgacett gaacatgaaa acaacaaaaa gggeetggte actgtagaag
                                                                       420
atgagcaggc gtggatggca tcttataaat atgtaggtgc tccactaata tccatncata
                                                                       480
tttgtccaca atgatcaact acgcccacct gtaaaggttc aaggttncat gtggcagaag
                                                                       540
aaccncatat tcattataca tggcttcttt tatgaatant cggccttggt tcttgaance
                                                                       600
cttgcaatga atttgnaatt ntacca
                                                                       626
      <210> 200
      <211> 618
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(618)
      <223> n = A,T,C or G
      <400> 200
actcataaaa aaagtcttac cccaaaattg caaacaaata cattaaaaga ttagaagagg
                                                                        60
tgacagaaag caccagacat taaacaaaat aaaaataata aaataaattc aactcaaaag
                                                                       120
gtccccattc agcaaatact ttgtaaagta tggcctgtat gtaaatagtg ctaaatcaag
                                                                       180
gactttttag cagaaaattg ctcggttctt ttatctaagg cttgaatttg taaagtgaaq
                                                                       240
gcataaaagt taccaaacat taagtaactc ttaaaatggc acacaggttt taaagctatt
                                                                       300
```

```
360
qqtttttcct tcctaactct ctgaattttt cccatggcct ttgtagatca actatttcaa
acqtatttta caccaqcaac totcaacata ottgtottto agatatgtca toagtcatgt
                                                                        420
ctaacaqqca aataqcanaa taacaqattt aaaacaatcc ttaactanct agcaqqacat
                                                                        480
                                                                        540
ttacttigga ttctgcataa ctgcaaactg acatatttgt aaagctaaaa atcagtttaa
                                                                        600
tentgattae agaaacteta teatgeteat taettaaeta ttgneettea ategetattn
                                                                        618
aaattcactt aatccaat
      <210> 201
      <211> 627
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(627)
      <223> n = A, T, C \text{ or } G
      <400> 201
                                                                         60
ggtactaggc acaatagaac atacagaaaa cattgtccct gctcttgagg agcttacatt
ctaaaagaaa aaatacacct tttttaaaaat ggcatttttg tttggtgttt tctgcaaagt
                                                                        120
acgcggggct ttttcttttt gaggaagacg cggtcgtaag ggctgaggat ttttggtccg cacgctcctg ctcctgactc accgctgttc gctctcgccg aggaacaagt cggtcaggaa
                                                                        180
                                                                        240
gcccgcncgc aacagccatg gcttttaagg ataccggaaa aacacccgtg gagtcggagg
                                                                        300
tggcaattca ccgaattcga atcaccctaa caagccgcan cgtaaaatcc ttggaaaagg
                                                                        360
tgtgtgctga cttgataaga ggcncanaag aaaagaatct canagtgaaa ggaccaagtt
                                                                        420
ngaatgccta ccaagacttt gagaatnact acgaganaaa ctccttgtgg tgaaggtcta
                                                                        480
agacgtgggn tngnttccag atgagaattc acaagcgact tattgacttc acaagtcctt
                                                                        540
ntgagattgt tangctgatt actteettna ntatganeen ngaatttaag ngggangtna
                                                                        600
                                                                        627
contnoagan gnttagttna ctatttt
      <210> 202
      <211> 620
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(620)
      <223> n = A, T, C or G
      <400> 202
actgcttaac gaaacactat cagcttgttt taaatggatc ttttaaatat caactgtagc
                                                                         60
                                                                        120
ctggttggct aattettet aatetteece attacttteg cetagattte ceatagatea
acaggcataq taaaatqcct catcaqaaca cacttctcca cacaattcaa aaaqggagct
                                                                        180
cctgtgggct caaagcaacc atcagtccaq caatqcccat qatttatctq aaactgcttc
                                                                        240
                                                                        300
ccaagagaca ggagtgcaga tetgagtage tgtgctgcca atacagatag gtttagcact
agatatttag tgattgtggc aaggaagaat cggtgatgat gggggtggtg ggtgaaggaa
                                                                        360
gggccagggg atctgaagga tcttcagttg ccttctcctg cttcttcatc ctgctggtcg
                                                                        420
480
                                                                        540
agcangettg etcangtgea ttetggatet catagtagaa caceggagaa ntganggeea
                                                                        600
ggcccaaccq qatnqqatqc
                                                                        620
```

```
<210> 203
      <211> 577
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (577)
      <223> n = A, T, C \text{ or } G
      <400> 203
60
atggaggcca tggggttggc ttgaaaccag ctttgggggg ttcgattcct tccttttttg
tctaaatttt atgtatacgg gttcttcnaa tgtgtggtag ggtgggggc atccatatag
tcactccagg tttatggagg gttcttctac tattaggact tttcgcttcn aagcgaaggc
                                                                             120
                                                                             180
                                                                             240
ttctcaaatc atqaaaatta ttaatattac tqctqttaqa naaatqaatq ancctacaqa
                                                                             300
tgataggatg tttcatgtgg ggtatgcatc ggggtantcc gagtaacgtc ggggcattcc
                                                                             360
                                                                             420
ggataggccn agaaagtgtt ntgggaanaa agttagattt accccgatga atatgatagt
gaaatggatt ttggcgtagg tttggtctag ggtgtancct gagaataggg gaaatccgtg
                                                                             480
aatgaaacct cctatgatgg caaatacact cctattgnta ggacataatg ngaagtgagc tacaaccgta atacctgccc nggcnggccc ttannan
                                                                             540
                                                                             577
       <210> 204
       <211> 629
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (629)
       <223> n = A, T, C or G
       <400> 204
cgaggtactt gtttttttt tttttttga gacggagtct cagtctgtca cccaggctag
                                                                              60
agtgcagtgg cacgacatcg gctcactgca acctccgcct cccgggttca agtgattctc
                                                                             120
ctgcctcaac ctcccgagta gctgggacta caggcatgtg ccaccacgcc tgactaattt
                                                                             180
ttgtattttt agtanagatg ggatttcatt atgttggcca gctggtcttg aacttctgag
                                                                             240
ctcaggtgat ccacccgcct tagcctncca gagtgctagg ataacaggca tgagccgtcg
                                                                             300
cgcctggcca aaatagcata atgttttaag aaagtttacg aatttgtctt gggccacatt
                                                                             360
naaaaccatc atgggccaag ggttggacaa gctagcctta ggtcatgtca gaatgcaatt
                                                                             420
taacaggaat ttcaagcnaa acttacaaaa aattaaatcc acaaaaaaaa tatcatttgg
                                                                             480
                                                                             540
taaatgcact gnctacacac tttactncta agtccattca accatgacga ccctttacat
aaaaattagg gcattctccc aagttctaaa gatgatttct aaaacattac caangnctaa
                                                                             600
agtctaattc ccacaaanca tttttttn
                                                                             629
       <210> 205
       <211> 424
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (424)
```

```
<223> n = A, T, C or G
       <400> 205
ggtacaaatg cttttatatt cagcccctgt aaagccatca gatgtttgaa agtttttaaa
                                                                             60
cacgaaccaa agggtttaat tttaagaact tagctaggaa tgggtgaaat cctacccaat
                                                                             120
taatagagtt ctgcaaatta gtaacaaagt gtaaaatgaa aggaagggtc ccttggagat
                                                                             180
gtgaaattet tetattgaga gteetgtett etttatteaa gaagtitgta geeatttea
                                                                            240
gaattcactc aagaaccaac ttcttaattt agatatcagc gaacaagtca tggcaaaaaa
                                                                            300
tacacaaaga gaaacaccac cacatcgaaa aggatgaaaa gccagaggtc caaccagtan
                                                                            360
gagtgtttgg gaagcccatt tgccccagac tgaggcctca catcgaagtt ctgcctcccc
                                                                            420
gcgt
                                                                            424
      <210> 206
       <211> 633
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (633)
      <223> n = A, T, C or G
      <400> 206
ggtaccaatg gtgcctcctg gaatcaagta tctttacctt aggaataacc agattgacca
                                                                             60
tattgatgaa aaggeetttg agaatgtaac tgatetgeag tggeteatte tagateacaa
                                                                            120
ccttctagaa aactccaaga taaaagggag agttttctct aaattgaaac aactgaagaa
                                                                            180
gctgcatata aaccacaaca acctgacaga gtctgtgggc ccacttccca aatctctgga
                                                                            240
ggatctgcag cttactcata acaagatcac aaagctgggc tcttttgaag gattggtaaa
                                                                            300
cetgacette atecatetee ageacaateg getgaaagag gatgetgtt cagetgettt taaaggtett aaateacteg aatacettga ettgagette aateagatag ceagactgee
                                                                            360
                                                                            420
ttctggtctc cctgtctctc ttctaactct ctacttagac aacaataaga tcagcaacat
                                                                            480
ccctgatgaa gtatttcaag cgtttaatgc tttgcagtat ctgcgtttat ctcacaacqa
                                                                            540
actggctgat agtggaatac ctggaaattc tttcaatggn gccatcctgg gtgaacctgg
                                                                            600
acttgcctat accagentaa aacataceae egg
                                                                            633
      <210> 207
      <211> 623
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (623)
      <223> n = A, T, C or G
      <400> 207
ggtacttttt ttttttttt tttttttt ttagaaacta tggctcttta ttttcatgtg
                                                                             60
gataattcaa acaaagtcat tagtagtctt tgttcaattt ttttttaaaa aacaaaaaaa
                                                                            120
ccctcaaata aaaaatcttg ggcttaaaag aactctatca caggagcctg gttggaggat
                                                                            180
tcctagtttt atacatgaga aatagaatgc agatttctct gaagagtgtt taaagaagga atggtagttg agggggctta tttcccaggc tcaaagtgat ttaggggtgg tgtcacagtg
                                                                            240
                                                                            300
ctaggtatag ggtgatggac agtgatcact gccgagggcc ttggaacgga tcttgctgtc
                                                                            360
acacaatgca ggtaacagag agtgggacaa caaaaagtaa tcaaggcgcc aaccaacatt
```

420

```
cttggatcga gcattcatat ataagtccaa aaggtgtang cataaggtgt gttggggtan
                                                                        480
aaqtqcctaa aqctqcaacc aqtgqcacan cctqcaqtaa ttccccqaac cttqqccttt
                                                                       540
tggggcgtga ancenceatt ettttggtne cetnggggtg enaaggeaat ttttnatqtq
                                                                        600
cccattgagg gttcaaacac aca
                                                                        623
      <210> 208
      <211> 620
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(620)
      \langle 223 \rangle n = A,T,C or G
acgatgtcta gtgatgagtt tgctaataca atgccagtca ggccacctac ggtgaaaaga
                                                                         60
aagatgaatc ctagggctca gagcactgca gcagatcatt tcatattgct tccqtqqaqt
                                                                        120
gtggcgagtc agctaaatac tttgacgccg gtggggatag cgatgattat ggtagcggag
                                                                        180
gtgaaatatg ccccgcgtac ttgctttgaa agattaccta ctattttatg ataaaatgta
                                                                        240
gttgtctcca gagcttaaat ataatttgta aagcacttgg tttaaatttc tctctaccta
                                                                        300
taaacagttt agcattaagg gtttctatta atgacacaga attattggcc aagtgtaatt
                                                                        360
tottaaaatt tagcattact ttaaatagco agcatgtaat acaagtaact acactaccto
                                                                        420
atatctacat gattttcaag ttgtaatgca gatggacaga taaaaaagat ttacqttqnc
                                                                        480
ttttggccat aagtgggaaa agttttctgn atattgcata gcattacaca tttatgccta
                                                                        540
ttttacatta actictaaag aagtttttct aagaaaangg ttcaggcaat attttttgag
                                                                        600
gctgccgaan aaaaatgant
                                                                        620
      <210> 209
      <211> 624
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(624)
      <223> n = A,T,C or G
      <400> 209
ggtactggta caaaaacagg cacataaacc aatgaaacag aatagaaagc ccagaaataa
                                                                         60
tgcttcaccc ccacaaccat ctgatcttca acaaaataaa caaaaacgag ccatggggaa
                                                                        120
aggactccct attcaataaa tggtgctggg ataactagtt aaccatatgc agaagattaa
                                                                        180
agetggaece etteettaca aaataaggag etggaeceet tatacaaaaa teaacteaaq
                                                                        240
atggattaaa gccttaaatg tgaaactata aaaccctgga agacaacata ggcgattcca
                                                                        300
ttctagacat cagaactggc aaagatttca tgaggaagac accaaaagca attgcaacaa
                                                                        360
aagcaaaaat tgacaactgg gatataatta agtttaagag cttctgcaca gcaaaagaga
                                                                        420
gactatcage agagtaaaca gaccacctac agaatgggag aaaatatttg caaactatge
                                                                        480
atgtgacaaa ggtctaatat ctagcatcta taagtactta aacaaatttc aacagaaaac
                                                                        540
caacacccca ttaaaaagtg ggcaaggaca tgaacaaatg cctttcaaaa gaagacatct
                                                                        600
gcttntacag tttntgaaac aaag
                                                                        624
      <210> 210
      <211> 504
```

```
<212> DNA
       <213> Homo sapiens
       <400> 210
acgcggggca gctagcagat gctttaggac ctagtatctg catgctgaag actcatgtag
                                                                           60
atattttgaa tgattttact ctggatgtga tgaaggagtt gataactctg gcaaaatgcc
                                                                          120
atgagttett gatatttgaa gaccggaagt ttgcagatat aggaaacaca gtgaaaaagc
                                                                          180
agtatgaagg aggtatettt aaaatagett eetgggeaga tetagtaaat geteaegtgg
                                                                          240
tgccaggete aggagttgtg aaaggeetge aagaagtggg cetgeetttg categggggt
                                                                          300
gcctccttat tgcggaaatg agctccaccg gctccctggc cactggggac tacactagag
                                                                          360
cageggttag aatggetgag gageactetg aatttgttgt tggttttatt tetggeteee
                                                                          420
gagtaagcat gaaaccagaa tttcttcact tgactccagg agttcagttg gaagcaggag
                                                                          480
gagataatct tggccaacag tacc
                                                                          504
      <210> 211
      <211> 619
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(619)
      <223> n = A,T,C or G
      <400> 211
accatgaaat atccagaaca tacttatatg taaagtatta tttatttgaa tccacaaaaa
                                                                           60
acaacaaata atttttaaat ataaggattt teetagatat tgeaegggag aatatacaaa
                                                                          120
tagcaaaatt gaggccaagg gccaagagaa tatccgaact ttaatttcag gaattgaatg ggtttgctag aatgtgatat ttgaagcatc acataaaaat gatgggacaa taaattttgc
                                                                          180
                                                                          240
cataaagtca aatttagetg gaaateetgg atttttttet gttaaatetg geaaceetag tetgetagee aggateeaca agteettgtt ceaetgtgee ttggtttete etttatttet
                                                                          300
                                                                          360
aagtggaaaa agtattagee accatettae etcacagtga tgttgtgagg acatgtggaa
                                                                          420
gcactttaag ttttttcatc ataacataaa ttattttcaa gtgtaactta ttaacctatt
                                                                         480
tattatttat gnatttattt aagcatcaaa tatttgtgca agaatttgga aaaatagaag
                                                                         540
600
ngangtttat taganaaan
                                                                         619
      <210> 212
      <211> 479
      <212> DNA
      <213> Homo sapiens
      <400> 212
cgaggtacaa agcagcaact gcaatactca aggttaaaac attagaaaag catttgtgtg
                                                                           60
acaggtatat tacagtatta tcaaaatatt acattttcag acttacttag cagataatca
                                                                         120
tccaccagag cttaaatctt taaattattt ccatagtctt aaaaaatatg taatgtcaga
                                                                         180
atgcatataa aaagaatgta aaaggaaacc taaaatacaa atggaataat gtaacaaata
                                                                         240
aatatttgat ttcagtaact gttaataatc agctcaacac caccattctc tctaaactca
                                                                         300
atttaattct tataggaata atgaactgtc aaatgccatg gcataattat ttatttccaa
                                                                         360
gctatcatca atgattagaa ctaaaaaaat tttggcataa aaaaatcaca attcagcata
                                                                         420
aataaagcta tttttagctt caacactagc tagcatctct aagaattgtt gaaataagt
                                                                         479
```

<210> 213

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<211> 487
      <212> DNA
      <213> Homo sapiens
      <400> 213
actittact geotogicae tatactitet atgeagatet cettititigigg tittecageet
                                                                           60
gteettteat cagageacat ggeageettt ggggtetttg gtetetgeea gateeatgee
                                                                          120
tttgtggatt acctgcgcag caagttgaat ccacaacaat ttgaagttct tttccggagc
                                                                          180
gtcatctctc tggtaggctt tgtccttctc accgtgggag ctctcctcat gctgacagga
                                                                          240
aaaatatete eetggaeggg gegtttetae teaetgetgg atecetetta tgetaagaae
                                                                          300
aacatcccca tcattgcttc tgtgtctgag catcagccca caacctggtc ctcatactat
                                                                          360
tttgacetge ageteetegt etteatgttt ceagttggee tetattactg etttageaac
                                                                          420
etgictgaig eceggattit tateateatg tatggtgtga ecageatgta ecteggeege
                                                                          480
gacacgc
                                                                          487
      <210> 214
      <211> 393
      <212> DNA
      <213> Homo sapiens
      <400> 214
cgaggtacaa tatgctgcag cataatttgt caggccaacc ttcacaccat attttqqcaq
                                                                           60
ttegtgtgca taegetgege agactateat atececetet ataegggcat aageaatetg
                                                                          120
acaaatgata tetetgtttg teacaegaae tateateetg tattigggtg tgttgtattt
                                                                          180
atttttatet tgtateacea agegttteeg ageataataa teagttttae cetetegteg
                                                                          240
tottotaaat ticactiggt atcicttaaa glaggootta ticttaacaa cittaacaaa
                                                                          300
ccccatectg cggaacagag accggcgtcc gctgctcgac agagacctgc aggcccagcg
                                                                          360
gcgctagggg gtgggaaaag ggccacccc cgt
                                                                          393
      <210> 215
      <211> 615
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(615)
      \langle 223 \rangle n = A,T,C or G
      <400> 215
ggtacagtaa caagtgttgg cattatcagt tgaactgtaa atacaaaatg cttcttccaa
                                                                           60
ttagteteta tgatgattaa qtttetaaaa tttatetgaa caccatteag aaaettgttt
                                                                          120
tggggaattt gatagttatt gatgtgcatc tgttaaactg atgacagaca taactcatca
                                                                          180
ttccccagaa acctttttg attacagtat ctaacatttt gcctcctctt ttttggtttt
                                                                          240
gctggttata aaggtttgga ttggagaggg ctcactggat cccaatcett ggagctggat cattggattc aaatcataat gtggatagga tagggaggat gaattaccag gattcatgga
                                                                          300
                                                                          360
gcgggatcag attaccagga acataggagt ggattcctgc ccaaccaaac ccqcattcqt
                                                                          420
gtggattttt ttattcaact taattggcta ttccaaagat tttttttcc tatttttqac
                                                                          480
gaatggagcc cttaagatgc acgatggaat tgggtttgcg tttttqqtaa aaqqaccaaa
                                                                          540
ccaggcctgg agataacgct ggagcaatct cntggaagga ttaqccccaa ttqatqqqaa
                                                                          600
catttaangg ggaag
                                                                          615
```

<210> 216

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<211> -322
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (322)
       <223> n = A, T, C \text{ or } G
       <400> 216
 ggtacttttt tttttttt tttttttt ttttttggag ttgtaggcaa atgtttaatt
                                                                             60
 aattetgete atatgeacat etgaaageat gagacacaet ceacagacag caegeactgg
                                                                            120
ggctggtggg gcanatgggc actcgccgat taggtattaa tgtcaataat acgtgcataa
                                                                            180
agtgctgata aaataactta agtgttacaa aaagagacag tccacggtgg ctgcaggcac
                                                                            240
atgcaggegg gactgggtca aacactccag ggctgcacat gttccagctg gcctgagtcc
                                                                            300
gacacgtcat aactggcctt gt
                                                                            322
       <210> 217
       <211> 606
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (606)
       <223> n = A, T, C or G
       <400> 217
acgcgggggg aagtgagcga cacactctgc gtcctcgcct caccagagtc ttgctgtgtg
                                                                            60
geccaggetg gagtgeegg etggteteaa atteetgace teaagtgate teeeteecaa
                                                                           120
agtgttgcga ttgcaggtgt gagccactgc acctggctgc tgagaaatct ttgcctacag tgagggaaac tactaaagtt cctggggaag caaagtaaga atttcataag aacaaaatgg
                                                                           180
                                                                           240
atggagagga gaaaacctat ggtggctgtg aaggacctga tgccatgtat gtcaaattga
                                                                           300
tatcatctga tggccatgaa tttattgtaa aaagagaaca tgcattaaca tcaggcacga
                                                                           360
taaaagccat gttgagtggc ccaagtcaat ttgctganaa cgaaaccaat gaggncaatt
                                                                           420
ttagagagat cottcacatg tgctatcgaa agtattcatg nattttacgt accttgggcc
                                                                           480
gegaceacet taaggeeaat theacacact ggenggeegt actantggat cenaetngga
                                                                           540
ccaacttggc gtaatcatgg catactggtt cctggggaaa atgtatccgt tacaattcnc
                                                                           600
acacan
                                                                           606
      <210> 218
      <211> 618
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (618)
      <223> n = A,T,C or G
      <400> 218
ggtacttttt ttttttttt tttttttga gacggagttt ggcccttgtt gcccaggctg
                                                                            60
aagtgcaata gtgcgatctc ggctcactgc aacctccacc ttccgtgttc aaccgattct
                                                                           120
```

```
cctgcctcag cctcctgagt agctgggatt acagatgaaa aaacatttaa agcccttaag
                                                                          180
qaaqaaqqaa atcaatqtqt aaatqacaaa aactataaaq acqccctcaq taaatacaqc
                                                                          240
gaatgettaa agattaacaa taaggaatgt gecatatata caaacagage tetetgttac
                                                                          300
ttgaagetgt geeagtttga agaageaaag caggaetgtg ateaggeact teagetaget
                                                                          360
gatgggaacg tgaaagcett ctatagacga actetggete ataaaggaet caagaattat cagaaaaget taattgatet caataaagtt ateetactag ateeaagtat tattgaggea
                                                                          420
                                                                          480
aagatggaac tggaagangt aactagactc ctaatcttaa ggataagaca gcaccattca
                                                                          540
acaaagaaaa ggagagaagg aaaatgagaa tcaagaggng aatgaaggca ngaggancet
                                                                          600
ggaaaacctg aggggagg
                                                                          618
      <210> 219
      <211> 613
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (613)
      <223> n = A,T,C or G
      <400> 219
ggtacaaagc ggatctgagc ccggaaaatg ctaagctcct cagcacattc ctaaatcaga
                                                                           60
ctggcctaga cgccttcctg ctagagctgc acgaaatgat aatcttgaaa ctaaagaacc
                                                                          120
eccaaaccca aacegaggag cgetteegee etcagtggag cetgagagae actetegtaa
                                                                          180
gttacatgca aactaaagaa agtgaaattc ttcctgaaat ggtatctcag ttcccagaag
                                                                          240
agatactget egecagetgt gteteagtgt ggaaaacage tgetgtgetg aaatggaate
                                                                          300
gagaaatgag atagaattat ttcctcagct atctttggat gactttggag agaagactcc
                                                                          360
teteteeteg tetgeggegt ggaettgate atggaetggt geetttgeat teagaaggag
                                                                          420
agotytoago ytagoacoga attoaaqaco aaqqoqtqot acctqaqotq acaqottttt
                                                                          480
gaaagccgag ctggttctga accatgtcct gcccnggcng gcgctcgaaa gggcgaattc
                                                                          540
agccactggc ggccgtacta ntggatccga actcggacca aacttggcgt aatatgggca
                                                                          600
tactggttcc tgg
                                                                          613
      <210> 220
      <211> 616
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(616)
      <223> n = A, T, C \text{ or } G
      <400> 220
ggtacgcggg ggcagccgcg gtgttgtgct gtgggggaagg gagaaggatt tgtaaacccc
                                                                           60
ggagcgaggt tetgettace cgaggceget getgtgegga gacceeeggg tgaagceace
                                                                          120
gtcatcatgt ctgaccagga ggcaaaacct tcaactgagg acttggggga taagaaggaa
                                                                          180
ggtgaatata ttaaactcaa agtcattgga caggatagca gtgagattca cttcaaagtg
                                                                          240
aaaatgacaa cacatctcaa gaaactcaaa gaatcatact gtcaaagaca gggtgttcca
                                                                          300
atgaattcac tcaggtttct ctttgagggt cagagaattg ctgataatca tactccaaaa
                                                                          360
gaactgggaa tggaggaaga agatgtgatt gaaagtttat cangaacaaa ccgggggtca
                                                                          420
ttcaacagtt tanatattot ttttaatnnt ttottttncc tcaatcottt tttattttta
                                                                          480
aaaatagttc ttttgtaatg tggtgtcaaa acggaattga aaactggcac cccatctttt
                                                                          540
```

```
gaaacatctg gtaatttgaa tctaatgctc attatcatta tggttggttt cattggcnga
                                                                        600
attttgggga tcaanc
                                                                        616
      <210> 221
      <211> 615
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(615)
      <223> n = A, T, C \text{ or } G
      <400> 221
ggtacagtga tagctccccc tgggcaatac aatacaagaa cagtgggttt tgtcaaattg
                                                                         60
gaacaaggaa acagaaccac agaaataaat acattggtta acatcagatt agttcaggtt
                                                                        120
actitititgt aaaagitaaa gtagagggga citcigtatt atqctaactc aaqtagactq
                                                                        180
gaatctcctg tgttcttttt tttttaaatt ggttttaatt ttttttaatt ggatctatct
                                                                        240
tetteettaa cattteagtt ggagtatgta geatttagea ceaetggete aatgegetea
                                                                        300
cctaggtgag agtgtgacca aatcttaaag cattagtgct attatcagtt accaccattt
                                                                        360
ggggctttta tectteatgg gttatgatge teteetgatg acacatttet etqaqttttq
                                                                        420
taattccagc caaagagaga ccattcacta tttgatggct ggctgcatgc agacatttaa
                                                                        480
agettttaga gaatacacta caccagggag tatgactact antatgacta ttagganggt
                                                                        540
aatacccaga attggactcg caccttaggc aagatccaac cactaaattg aataagaatg
                                                                        600
agtngatgag gtncc
                                                                        615
      <210> 222
      <211> 617
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(617)
      <223> n = A, T, C \text{ or } G
      <400> 222
ggtacttttt ttttttttt tttttttt ttttaattta tgattttatt gnctttcctt
                                                                         60
tgtccggcct ttaacatgtt tctgtaattt aaataaaaat ctatttactt tctccatttt
                                                                        120
agcaaatggt ttetttaeee aaataggttg eactatagte eecatatggt tttetaetgn
                                                                        180
tccacaacca ctatttcaca aagattgaca aaactttaat aaaagttaaa tttacagaca
                                                                        240
tcttaagata acttgggaaa tatgtagtaa aaaagaatcg agtccacaaa ttaagaatat
                                                                        300
tttgctaata tgcccaacac caatttcagc aaatccaatc tacttaactc atatatttaa
                                                                        360
tgnggtaatt tttctaacaa aatttaatgg gggtatgaat gatatattta tqcccttqac
                                                                        420
aaagatgaca tgtgtgattt tggtgngact aanaaaggag aagtatgatt tctggngggt
                                                                        480
atganatcac tetggeteat egaageteea gaatatgtaa gggtetgnea egteeaaaaa
                                                                        540
tgttaggcna atgtataaaa ggccacccgg ctnacacacg ttttatatac aaactttngn
                                                                        600
agtcctttta tntcata
                                                                        617
      <210> 223
      <211> 470
      <212> DNA
      <213> Homo sapiens
```

```
<220>
      <221> misc feature
      <222> (1) ... (470)
      <223> n = A, T, C \text{ or } G
      <400> 223
ggtaccacaa ctgtgccctt gataattagt aatcactcct aaaaatcttc atttggcacc
                                                                            60
agatggtgtg tttaaaacac cctaggatgt tttgaatcag gcttgatttt gttagttgag
                                                                           120
ttacaggaga attttaaggg tgagggtatg ggggtcaggg aagaaaagga aatgggaaat
                                                                           180
ggaccagaaa aaatettgag teateateta aateaacaaa geactgatag etecaaatat
                                                                           240
taggtcagac actaaaacga ctgatatagg ctcaagtggt ttataaaaacc tataaaaaqa
                                                                           300
ctacaccage aaagteeetg teaatetgte agagtteaga aactaaaaca gggagtaaca
                                                                           360
ttttagctta aaaccttatc tcaagagaat catatacact tcacatgaat aaaaatacct
                                                                           420
gaaaccaaac atttttaaaa geteeagtee tgeeenggee ggeegetega
                                                                           470
      <210> 224
      <211> 622
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (622)
      <223> n = A,T,C or G
      <400> 224
gcgtggncgc ggccgacgtn ctctttttt ttttttttt tttttttgcnn actaaaaatn
                                                                            60
ngattgetet ttaaageett aggeegnatg acaaaatgan nagaetgaaa tgacaneggg
                                                                           120
gaggaagaaa cagannaaag ataagaatga ggtggtcagg ttgggggaat taagcgaata
                                                                           180
ttenetteen nggtgagtee tnacactggt eteatgeeca tgatgagttg cacaccaaac
                                                                           240
acnggetgnt gactinecte etgenetant cagtgaacti gengacatng ggnanectea
                                                                           300
cattacagnt ataanntttc cacctaaaaa atgctgcgct tttcgacngg ctcnnncagn
                                                                           360
ggccggggct tgacatggng gaanggattt ctctcccatg ccaaggaatt catcacatca ctgntactcc actgncaacc ttntccattg ggctcngtgc cctgtgtngg gtcatggacc
                                                                           420
                                                                           480
cantccanaa ntatgaatac tgtaccatgc tcttaaccag gaggacctaa ggatccttag
                                                                           540
nccentgagn nanacaccag gnttcaaagg cegttttggn aagccaaatt tgnttnggne
                                                                           600
cgaattnggg ccaaacangg tt
                                                                           622
      <210> 225
      <211> 619
      <212> DNA
     <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (619)
      <223> n = A, T, C \text{ or } G
      <400> 225
acgcggggag ttccgccatg gcctccttgg aagtcagtcg tagtcctcgc aggtctcggc
                                                                            60
gggagctgga agtgcgcagt ccacgacaga acaaatattc ggtgctttta cctacctaca
                                                                           120
acgagegega gaacetgeeg eteategtgt ggetgetggt gaaaagette teegagagtg
                                                                           180
```

```
gaatcaacta tgaaattata atcatagatg atggaagccc agatggaaca agggatgttg
                                                                            240
ctgaacagtt ggagaagatc tatgggtcag acagaattct tctaagacca cgagagaaaa
                                                                            300
agttgggact aggaactgca tatattcatg gaatgaaaca tgccacaqqa aactacatca
                                                                            360
ttattatgga tgctgatctc tcacaccatc caaaatttat tcctgaattt attagcccgt
                                                                            420
ggggccaatt ttttaactca natcttgctg agaccaggag catctgattt aacaggaagt
                                                                            480
ttcagattat acccgaaaaa gaagttctag agaaattaat agaaaaatgt ggttctaaag gctacgtctt ncaaatggag atgattggtc nggcaagaca gttgaatatt ctattggcga
                                                                            540
                                                                            600
ggttccatat canttgngg
                                                                            619
      <210> 226
      <211> 277
       <212> DNA
      <213> Homo sapiens
      <400> 226
acgcggggcc cctcatttac ataaatatta tactagcatt taccatctca cttctaggaa
                                                                             60
tactagtata tegeteacae eteatateet ecetaetatg cetagaagga ataatactat
                                                                            120
cgctgttcat tatagctact ctcataaccc tcaacaccca ctccctctta gccaatattg
                                                                            180
tgcctattgc catactagtc tttgccgcct gcgaagcagc ggggggccta gccctactag
                                                                            240
tctcaatctc caacacatat ggcctagact acgtacc
                                                                            277
      <210> 227
      <211> 328
      <212> DNA
      <213> Homo sapiens
      <400> 227
ggtacatatt tttgccaatg ctatacagca aaaatgaaaa acttacagaa aggtaaacaa
                                                                             60
aattgagtcc acttttttaa tttcacaagc tgctttaaac tatagaacca ccagatatct
                                                                            120
gtaaaataag caaaactggt aagtgtgttt tittaattga gggaaggagg gccagaggag
                                                                            180
ttggtgcaga agcgcttcgg gtgaattcat accagagcca ccgggtgtga ctcggctacc
                                                                            240
tctcccaatt accacaggga ggtcttaaaa ttgaatttca gtttcagcag atactccaga
                                                                            300
tttacctgag caatatcata gacaatgt
                                                                            328
      <210> 228
      <211> 609
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(609)
      <223> n = A, T, C or G
      <400> 228
acgcgggagt tcaagcagat gtatggctaa ccggaaacag gtgggtcacc tcctgcaaga
                                                                             60
agtggggcct cgagctgtca gtcatcatgg tgctatcctc tgaacccctc agctgccact
                                                                            120
gcaacagtgg gcttaagggt gtctgagcag gagaggaaag ataagctctt cgtggtgccc
                                                                            180
acgatgetca ggtttggtaa ecegggagtg tteecaggtg geettagaaa geaaagettg taaetggeaa gggatgatgt cagatteage ecaaggttee teeteteeta eeaageagga
                                                                            240
                                                                            300
ggccaggaac ttetttggac ttggaaggtg tgcggggact ggccgaggcc cctgcaccct
                                                                            360
gcgcatcagg actgcttcat cgtcttggct gagaaaggga aaagacacac aagtcgcgtg
                                                                            420
ggttggagaa gccagancca ttccacctcc cttccccaac atctctcana gatgtgaaac
                                                                            480
```

```
cagateteat ggeaacnaag ceetntgeaa gaageteaag gaanetaagg aaaatggaeg
                                                                       540
ttttcagana atggttgtag ttcatgggtt ttncctactg ccgggtcctt tcttangacc
                                                                       600
                                                                       609
cgcanaant
      <210> 229
      <211> 610
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(610)
      \langle 223 \rangle n = A,T,C or G
      <400> 229
ggtacttttt tttttttt ttttttttt gcagactaaa aattttattg ctctttaaaq
                                                                         60
ccttaggccg tatgacaaaa tgaagagact gaaatgacag cggggaggaa gaaacagaag
                                                                        120
aaagataaga atgaggtggt caggttgggg gaattaagcg aatattctct tccagggtga
                                                                        180
gtecteacae tggteteatg eccatgatga gttgeacaee aaacacagge tgetgaette
                                                                        240
ceteetgeae tagteagtga aettgeagae atagggtaae eteacattae agttataate
                                                                        300
tttecacete agaaatgetg tgettetega caggetegca cagtggeegg ggettganat
                                                                        360
ggtggaggga tttctctccc atgcaaagta attcatcaca tcactgntac tccactccca
                                                                        420
acctteteca ttgggetegg tgeeetgtgt ggggteatgg acceaateca acgtatgant
                                                                        480
actggtacca atgctnttac cagggaggac acnaaaggat cccttacccc ctgagcacag
                                                                        540
acconaggit tcaaanggco gittiggcag gccaaactqn aintqnccag aaittgqnqa
                                                                        600
caaaacaagg
                                                                        610
      <210> 230
      <211> 346
      <212> DNA
      <213> Homo sapiens
      <400> 230
ggtcggccga ggtaccatgc actgagtgac tgtggggatc atgttqttat aatgaacaca
                                                                         60
agacacattg cattttctgg aaacaaatgg gaacaaaaag tatactcttc gcatactggc
                                                                        120
tacccaggtg gatttagaca agtaacagct gctcagcttc acctgaggga tccagtggca
                                                                        180
attgtaaaac tagctattta tggcatgctg ccaaaaaacc ttcacagaag aacaatgatg
                                                                        240
gaaaggttgc atcttttcc agatgagtat attccagaag atattcttaa gaatttagta
                                                                        300
gaggagette etcaaceacg aaaaatacet aaacgtetag atgagt
                                                                        346
      <210> 231
      <211> 601
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(601)
      <223> n = A,T,C or G
ggtacgcggg gagagcacat ccggtgttag aagcgctggt aggccttgga gaggcgggtt
                                                                         60
aggaagagtg gagactgctg cacggactct ggaaccatga acatatttga tcgaaagatc
                                                                        120
```

```
aactttgatg cgcttttaaa attttctcat ataaccccgt caacgcagca gcacctgaag
                                                                                      180
aaggtetatg caagttttge cetttgtatg tttgtggegg etgeagggge etatgteeat atggteacte attteattea ggetggeetg etgtetgeet tgggeteeet gatattgatg atttggetga tggeaacace teatageeat gaaactgaac agaaaagact gggaettett
                                                                                      240
                                                                                      300
                                                                                      360
getggatttg catteettac aggagttggc etgggeeetg eetggagttt tgnattgetg
                                                                                      420
teaaccccae atcettecae tgettteatg ggeeegeaat gatetttace tgettaacet
                                                                                      480
taatgcactc tatccaagcg ccgtactcct tttctgggag gatcttgatg tcagcctgaa cttgtgcttt gcttcctggg gaatgtttct ttggatccat tggcttttca gcnaactttt
                                                                                      540
                                                                                      600
                                                                                      601
       <210> 232
       <211> 390
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (390)
       <223> n = A, T, C or G
       <400> 232
actititit tittitit tittitit tiggittiaa tgittatiic cccaagacag
                                                                                       60
cctagcctgc actctacttg gataaatttt acaagctagt tttctgctgc ttctagtttt
                                                                                      120
aaactttaac catgtttctg atgacaagga atgctgcaaa aatactctag ttcaacaaag agttatgatc acaaaataat ttttatccat tctacagtgt ttcanaatta ccagttgatt
                                                                                      180
                                                                                      240
tttaaacaca aagtagatat agatgctaat ggtggctaat ctggtatgtt tcttatagca
                                                                                      300
aactgttgtt catgcaacac ttgtgctcaa aggggaaggc acaggatttc ctacaatgag
                                                                                      360
ccaccttata aagagttctt tttqnacctn
                                                                                      390
       <210> 233
       <211> 603
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(603)
       <223> n = A,T,C or G
       <400> 233
cgaggtacgc gggggaagag tgagggttcc aacttttctg cttatctggg aggtgttggg
                                                                                       60
cgcggacaat cgagatgtca gagaaaaagc agccggtaga cttaggtctg ttagaggaag
                                                                                      120
acgacgagtt tgaagagttc cctgccgaag actgggctgg cttagatgaa gatgaagatg cacatgtctg ggaggataat tgggatgatg acaatgtaga ggatgacttc tctaatcagt
                                                                                      180
                                                                                      240
tacgagetga actagagaaa catggttata agatggagac ttcatagcat ccagaagaag
                                                                                      300
tgttgaagta acctaaactt gacctgctta atacattcta gggcagagaa cccaggatgg
                                                                                      360
gacactaaaa aaatgtgttt atttcattat ctgcttggat ttatttgtgt ttttgtaaca
                                                                                      420
caaaaaataa atggtttgat ataagaaaaa annnnnnnna aaaaaaaagt nctggccngg
                                                                                      480
eggeegttea aanggeeaat tecaceeact ggeggeegta etaanggaee aacttggnee
                                                                                      540
aacttgggga atcanggcaa actqqttcct qqnqaaatqq nttcccttcc aattcccaa
                                                                                      600
atn
                                                                                      603
```

```
<211> 616
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (616)
       <223> n = A, T, C or G
       <400> 234
cgaggtacct tcattgcgat caaaccagat ggggtccagc ggggtcttgt gggagagatt
                                                                                  60
                                                                                 120
atcaaqcqtt ttgagcagaa aggattccgc cttgttggtc tgaaattcat gcaagcttcc
gaagatette teaaggaaca etaegttgae etgaaggaee gteeattett tgeeggeetg
                                                                                 180
gtgaaataca tgcactcagg gccggtagtt gccatggtct gggaggggct gaatgtggtg
                                                                                 240
                                                                                 300
aaqacqqqcc qaqtcatqct cqgggagacc aaccctgcag actccaagcc tgggaccatc
                                                                                 360
cgtggagact tetgcataca agttggcagg aacattatac atggcagtga ttetgtggag
agtgcagaga aggagatcgg cttgtggttt caccetgagg aactggtaga ttacacgaac tgtgctcana actggatcta tgaatgacag gaaggcagac ccattgnttt tcacatneat ttccettent tecattggc aaaggaccag etttnggaaa tetantnttt acenggacet
                                                                                 420
                                                                                 480
                                                                                 540
tattettaat ttqqanqqaa actnttggae tttgangtnt teetntaeet ngeeegggng
                                                                                 600
gccgtttaaa agggna
                                                                                 616
       <210> 235
       <211> 607
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (607)
       <223> n = A, T, C or G
       <400> 235
acqcqqqqqq tqcqttactt acctcqactc ttaqcttqtc qqqqacqqta accqgqaccc
                                                                                  60
ggtgtetget cetgtegeet tegeeteeta atecetagee actatgegtg agtgeatete
                                                                                 120
catecaegtt ggecaggetg gtgtccagat tggcaatgee tgetgggage tetaetgeet
                                                                                 180
ggaacacggc atccagcccg atggccagat gccaagtgac aagaccattg ggggaggaga
                                                                                 240
                                                                                 300
tgactccttc aacaccttct tcagtgagac gggcgctggc aagcacgtgc cccgggctgt
gtttgtagac ttggaaccca cagtcattga tgaagttege actggcacct accgccagct
                                                                                 360
                                                                                 420
cttcaccctg agcagctcat cacaggcaag gaagatgctg ccaataacta tgcccgangg
cactacacca ttggcaagga gatcattgac cttgngttgg acccaattcc aaacctggct
                                                                                 480
gaccatgcac cgggctttan ggnttnttgg gttttcccaa antttggggg ggaactgggt ttgggttaac ttcctgntna tggnacgntt ttaaatgaat ntgggaaaaa tccaactggn
                                                                                 540
                                                                                 600
                                                                                 607
gntttcc
       <210> 236
       <211> 608
       <212> DNA
       <213> Homo sapiens
       <221> misc feature
       <222> (1) ... (608)
```

```
\langle 223 \rangle n = A, T, C \text{ or } G
      <400> 236
acgogggcat gcaacaccac acccagcctg aaacccagat ttttaatatg aaatcaaagt
                                                                          60
cttcagacct tgtaggtgtc ataaaaagca cgctgaggac cactagtttg caactgccaa
                                                                         120
tetaaaatat catagacatt atateaette aaceaegaaa aaaaagtatg tgaggeagaa
                                                                         180
aatggaagca accatgccta atttattgtt gaatactttt tccgtatacc aagagcttcc
                                                                         240
tttgcactag catctgaaac tatatccaga atgacactgg ttttcataaa agtgttgatc
                                                                         300
ctcacacctc tttatagtct tgcacctagc acagtqqaqt qaaacacttt aaataqcact
                                                                         360
tgntccttga gtatatatgg aaaaaagtga agtattgata aagtgctcaa ctaatatqaq
                                                                         420
cagcatetea ggagteteca attettgaat taccagggag tatttttace attttcccca
                                                                         480
ntgnaaggcc ttttttgaga nacttaccct caaatngaan gnnttaagca tqntcctttt
                                                                         540
tttttccttt ttttttgan aaaagggctt gctntgtggc caggttggan tgcctacntg
                                                                         600
aaaattcn
                                                                         608
      <210> 237
      <211> 609
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(609)
      \langle 223 \rangle n = A,T,C or G
      <400> 237
actatttcat atattgtgtg agccccacaa atgtctattt taaaaagagt atagtccctg
                                                                          60
gecaggegeg gtggeteacg cetgtaatec cagcagtttg ggaggeegag gtgggeggat
                                                                         120
cacctgaggt ctggagttcg agaccagcct gaccaatatg gtgaaacccc gtttctacta
                                                                         180
aaaatacaaa attagctggg catggtggag catgcctgta atcccagcta ctcgggaggc
                                                                         240
tgaggcagga gaatcacttg aaccegggag gegaaggetg cagtgageca agatcaegee
                                                                         300
attgcactcc agcctgagca acaagaggga cactccgtcc ccaaaaaaaa aataataaaa
                                                                         360
aaaataaaaa ataaaaataa aaagagtata gttcccaatg ggttctacaa acattcctga
                                                                         420
tttatactgg gggaagtgat gcctaantgg gaacattaat cattatggtt tcgaaaatta
                                                                         480
aatatttetg caaacaatte etttgeaaat getaaettge eatgagetta eeceatttga
                                                                         540
aattgngnet ttacaaagac cttggeegga ceeettangg ngaatteagn caetggnggg
                                                                         600
cgttctttg
                                                                         609
      <210> 238
      <211> 616
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (616)
      <223> n = A,T,C or G
acgaggcggt gcgggaagtc ctgcacggga accagcgcaa gcgccgcaag ttcctggaga
                                                                          60
cggtggagtt gcagatcagc ttgaagaact atgatcccca gaaggacaag cgcttctcgg
                                                                         120
gcaccgtcag gettaagtee acteceegee ctaagttete tgtgtgtgte etgggggace
                                                                         180
agcagcactg tgacgagget aaggcegtgg atateceeca catggacate gaggegetga
                                                                         240
```

```
300
aaaaactcaa caagaataaa aaactggtca agaagctggc caagaagtat gatgcgtttt
                                                                       360
tggcctcaga gtctctgatc aagcagattc cacgaatcct cggcccaggt ttaaataagg
caggaaaagt teeetteetg eteacacaca acqaaaacat ggtggecaaa agtggatgag
                                                                       420
                                                                       480
gtgaagtcca caatcaagtt ccaatgaaga aggggtatgt ctggcttgta acttgttggt
cacgtgaaga tgacngacga tgacttgngt ataacattna netgggetgg caacttettg
                                                                       540
                                                                       600
gggcaatgnt caanaaact ggcaaaatgt ccgggccttt tttttagagc cccttggnaa
accccangcc ntttta
                                                                       616
      <210> 239
      <211> 607
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(607)
      <223> n = A,T,C or G
      <400> 239
acagtetgtt egagaacace ttggteatga aagtgacaac etgetgtttg tteagateae
                                                                        60
aggcaaaaaa ccaaactttg aagtgggttc ttctaggcag cttaagcttt ccatcaccaa
                                                                       120
gaagtettet eetteagtga aacetgetgt ggaceetget getgecaage tgtggaceet
                                                                       180
ctcagccaac gatatggagg acgacagcat ggatctcatt gactcagatg agctgctgga
                                                                       240
tccagaagat ttgaagaagc cagatccagc ttccctgcgg gctgcttctt gtggggaaag
                                                                       300
ggaaaaagag gaaggcctgt aagaactgca cctgtggcct tgccgaagaa ctggaaaaag
                                                                       360
agaagtcaag ggaacagatg aacttccaac ccaagtcaac ttgtggaaac tgctcctggg
                                                                       420
cgatgcetti cgttgtgcca ctggccctac cttgggatgc cagcntnaaa ctggggaaaa
                                                                       480
gngcttctaa tgatancatc tttattgaag cctaagaagg ttctgaattg ggacccattt
                                                                       540
gttetteaac caattetggn ettaaateea eettgggggt etteeacete ettggatttg
                                                                       600
ncacctt
                                                                       607
      <210> 240
      <211> 615
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(615)
      <223> n = A,T,C or G
      <400> 240
ggtacgcggg gcttttcaca agatggcgcc gaaagcgaag aaggaagctc ctgcccctcc
                                                                        60
taaagetgaa gecaaagega aggetttaaa ggecaagaag geagtgttga aaggtgteea
                                                                       120
cagccacaaa aagaagaaga teegeaegte acceaeette eggeggeega agacaetgeg
                                                                       180
actooggaga cagoocaaat atootoggaa gagogotoco aggagaaaca agottgacca
                                                                       240
ctatgctatc atcaagtttc cgctgaccac tgagtctgcc atgaagaaga tagaagacaa
                                                                       300
caacacactt gtgttcattg tggatgttaa agccaacaag caccagatta aacaggctgt
                                                                       360
gaagaactgt atgacattga tgtggccaag gtcaacaccc tgattcggcc tgatggagag
                                                                       420
aagaaggcat atgttegact ggeteetgat tacnatgett tggatgttge caccaaaatt
                                                                       480
gggatcattt aactgagtcc acttgctaaa tctgaatata tatatata tatatctttt
                                                                       540
cnccccaaaa aaaaaaaaa aaaaaagtnc tncccggcgg ccgtttaaag gggaattccc
                                                                       600
cacttggggg cgttt
                                                                       615
```

```
<210> 241
      <211> 365
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (365)
      <223> n = A, T, C \text{ or } G
      <400> 241
acggggggt cgctttgctg ttcgtgatat gagacagaca gttgcggtgg gtgtcatcaa
agcagtggac aagaaggetg etggagetgg caaggteace aagtetgeec agaaagetea
                                                                        120
gaaggctaaa tgaatattat ccctaatacc tgccacccca ctcttaatca gtggtggaag
                                                                        180
aacggtctca gaactgtttg tttcaattgg ccatttaagt ttagtagtaa aagactggtt
                                                                        240
300
tggttttctt ttttgcgtgt ggcaagtttt aaagttatta agtttttaaa atcaagtacc
                                                                        360
tnggn
                                                                        365
      <210> 242
      <211> 625
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (625)
      <223> n = A, T, C or G
      <400> 242
natngganng nttttccctt aacgtgggcc ncggccgagg nactttttt tttttttt
                                                                        60
ttttttttt gcaggcagct atttaattan gntcttaana catttanaac nccaatttgn
                                                                        120
gaanataaat teeattegte anaacaaacn cagategean gtageeetgg anetgangaa
                                                                        180
taactttgat ttttggnaaa atttgngagt concagettt etgateaate ttgegetget
                                                                        240
cccnaatete atattetet ttttetgggg ccaaaatett acetteetgg ngtetggget ttegcaaett ettettettg aaagaageet cagtaaaaat ggtttgggaa ttttacatta
                                                                        300
                                                                        360
ctgatatcca atttnggtga aatggcaatg accaatttct ngggggggtct tcgtaaaaga
                                                                        420
actocantga nggnocaaag gtocagtoco aagtataggo notnaccact gnttcaggaa
                                                                        480
accacctttt gncctggggg gtccatgagg atgaccaaat ggncccgggg naagctggct
                                                                        540
ccantttttt acggcctacc gaagggtttt tgccngggta aaagttttaq qqccattttc
                                                                        600
ngggnaaatc taggcttttg gaaat
                                                                        625
      <210> 243
      <211> 639
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(639)
      <223> n = A, T, C or G
```

```
<400> 243
nncnaattcc nccntaaccn ggnccccgnc caagnacccc ggcncctttg gatgtatnga
                                                                             60
aatnaacnta ttaatgggga cntattggag aaggaaatno ctagacctac aactttnago
                                                                            120
naatagengt gatgttttag gaactgaaat gteacaetta aagtettnag eecagetaet
                                                                            180
tccctatttt tgtggggaga aaanggccng attagaactg ttctggttgt gtttggcggg
                                                                            240
aggggaataa tttttgttca gtccttctta gtgaccaaac tttaattttt aagaataata
                                                                            300
tattgactta ctgaactgaa gcattctgag ttgaaaggag ctccncagga ntggagttct
                                                                            360
gtgttgctca catgttnaaa nettgctcac ettnatagen caaggaatac etatetteca
                                                                            420
natneegeea tttteatete ttaaatgnag teeaaagtat gaettgagaa agttgetetn
                                                                            480
ggattctggg gtcttaaaac tngggattct gggattntgg ggtccnaaag ttnaccttgn
                                                                            540
aaagttgcct gggnttttan aaatnenetg nattetgggg ttttaaaaaa ttttgaaaaa
                                                                            600
acccencen nettgaaagg gacettaaaa attaacetn
                                                                            639
       <210> 244
       <211> 614
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
       <222> (1) ... (614)
       <223> n = A, T, C or G
      <400> 244
tegageegne ggeeeggee aggtaetttt ttttttttt tttttttt gaaaatggag
                                                                             60
tettgetetg ntgecaaact ggantgeaat ggtgeganet gggeteactg naateteeac
                                                                            120
ctnccgggtt caagcgattc tectgcetca cetecgagta actgggacta caggtgegeg
                                                                            180
ccaccaagcc cagctcattt ttgnattttt agtanaaatg gggtttcacg atgttggcta
                                                                            240
ngatggnete gatetetggt caaagtettt tetgnaaata teettggtaa aaaaacaatt
                                                                            300
ttagactgta gctgttgcaa atgctttaag gaagaaacna aacaactgca gtcttcctga aatgaaaaaa ctccccaggg ctgctattna aaacaacccc accagcactt caatcatgat
                                                                            360
                                                                            420
gecnacagtg geccaetgaa aaanenggaa aagttenaat eecaaaetgg gatgetettg
                                                                            480
actntggaat tntgngggen ntnecenant ttnanacaaa aengnetngg necetntttt
                                                                            540
ttgggggaat ttgggaanaa aaaaacttgn gngttcttgn ggttccnttg ttccccaaaa
                                                                            600
nactgggggn nggg
                                                                            614
      <210> 245
      <211> 620
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (620)
      <223> n = A, T, C \text{ or } G
      <400> 245
gccgtggtcg cgggccgagg tccatttgcc tcccggcctc aagccgattc tcctgcctca
                                                                             60
geoetecaag tagetgggga ttacaggeac etgecaceat geoeggetaa tttttgnaat
                                                                            120
tttagtagag acagggtttc accatgttgc ccaggctggt ttcgaactcc tgacctcagg
                                                                            180
tgatccaccc gcctcggcct ccaaagtgct gggattacag gcttgagccc ccgcgcccag ccatcaaaat gctttttatt tctgcatatg ttgaatactt tttacaattt aaaaaaatga
                                                                            240
                                                                            300
tctgntttga aggcaaaatt gcaaatcttg aaattaagaa ggcaaaaatg taaaggagtc
                                                                            360
```

```
aaaactataa atcaagtatt tgggaaagtg aagactggaa gctaatttgc attaaattca caaactttta tactctttct ggatatacat tttttttctt taaaaaacaa ctttngatca
                                                                                 420
                                                                                 480
gaatageee atttagaace ttttggtate agneaatatt tttaaatagt tnaacengge
                                                                                 540
ctaagetnaa aqnqqettga tntgagtaaa ettttcaact ggettgaace etnaacettt
                                                                                 600
taaaatgacc ttccgagntt
                                                                                 620
       <210> 246
       <211> 595
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(595)
      \langle 223 \rangle n = A,T,C or G
       <400> 246
acttattett caggggttac tgagteggea cetatgacag etaagagage tttettaaag
                                                                                  60
actgoctcag tgtcttcttg gcttttggca cottcactcc actctgcca ggaaatccac
                                                                                 120
aatggcagac aaacctgggg tttcaggtgc acaaagactt cttcaaaaag catggctatq
                                                                                 180
tcagggctct ttgactcgat cagcacctgc agcttcagct gccacattgt cccagagtct
                                                                                 240
ctaaacaatt caagttccag ctactgncac ttccagagct tcctcaggaa gttataacac
                                                                                 300
agcaacgaaa cactcaactg cttgtattgg cattctgaca gaagcttcaa gttcatgtgc cttcctgaat acagtcatgg tctttncaac ctcttcctct aaggacccac tatttgactt
                                                                                 360
                                                                                 420
cttaataaat ctttccagcc aaaggngatg aacactttca catgggcctt gtggcaaaag
                                                                                 480
cttnatggct ttttatcncg gacagacctt tctcttcggg cgacctcaat ggtttggctt
                                                                                 540
ggtcgtggag ctggtntttg gctnggactc aacttnaatn ttgcttqccc naaac
                                                                                 595
       <210> 247
       <211> 364
       <212> DNA
       <213> Homo sapiens
       <400> 247
gggtacacta gaaagtcttt tacaaaataa tcatcttaga tcaacagaag accaatcttc
                                                                                  60
aatgtcgtcc tgcaagatgg gttactttaa catctcctcc tgttttctcc aatgttctcc tttagtatgg ctggtaattg ttttggtgat tgccacccc tcgagatgcc ttgccataag
                                                                                 120
                                                                                 180
tgctctgttg gccactgtag tctgcatatc cctgtccata tccatagttc ccatagttat
                                                                                 240
acccagtata atcatatccg ccatagccac tatagttttg atcaccacca taggcactat
                                                                                 300
tgtaatttcc atatccttga tcataatagt tattaaatcc ttggttccag ttttggccct
                                                                                 360
qacc
                                                                                 364
       <210> 248
      <211> 591
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (591)
      <223> n = A,T,C or G
      <400> 248
```

```
ggtncagata tettcaaagg aggaagaaga aagggaaace agatggtqqa cetqaatatg
                                                                        60
ncccttance aganctaate aacccactea gecagaatag aagaagetgg aatagattee
                                                                       120
ccaacctggt ttgccagttc atcttttgac tctattaaaa tcttcaatag ttggtattct
                                                                       180
gnaatttcac teteatgant genactgngg ettaactaat attgcaatgn ggettgaatg
                                                                       240
taagtagcat cetttgatge ttetttgaaa ettgnatgaa tttgggtatg aacagattge
                                                                       300
ctgctttccc ttaaataaca cttaaaatta tttggaccag tcagcacaac atgcctnggt
                                                                       360
tgnattaaag ennggatatg etggatttta taaaattgge caaattagag aaatntagte
                                                                       420
ccatggaaat atattettg taaaaaagtg ettgaatett tttggtcaag ataatgecae
                                                                       480
tottaagaat atottoncac tnttgangga ttaaatatog goantggaaa agoottaaaa
                                                                       540
atggggtcna cttgccttgn gcctaaaccg accctgaaat gggatttccc n
                                                                       591
      <210> 249
      <211> 332
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (332)
      <223> n = A,T,C or G
      <400> 249
actotocgag agggtogttt tocogtocoo gagagcaagt ttatttacca aatgttggag
                                                                        60
taataaagaa aggcagaaca aaatgagctg ggctttggaa gaatggaaag aaagggctgc
                                                                       120
ctcaagaget ettcagaaaa ttcaagaaet tgaaaggaca gettgacaaa etgaagaagg
                                                                       180
aaaagcagca aaggcagttt cagctttgac agtctcgagg cttgcgcttg cagaaacnaa
                                                                       240
aacagaaagg ttgaaaatga aaaaacccag ggtaccttgg nccgggacca cgcttaaggc
                                                                       300
gaaattccaa cacacttggc cggccggtac ta
                                                                       332
      <210> 250
      <211> 626
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (626)
      <223> n = A, T, C or G
      <400> 250
ggtactatta gccatggtca accccaccgt gttcttcgac attgccgtcg accggcgaag
                                                                        60
ccctttggcc cgcgtcttcc tttgaactgg ttgcagacaa gggtcccaaa ganagcagaa
                                                                       120
aattttegtg etetgageae tggagaaaaa ggatttggtt ataagggtte etgettteae
                                                                       180
agaattattc cagggtttat gtgtcaaggt ggtgacttca cacgccataa tggcactggt
                                                                       240
ggcaaagtcc atctatgggg aagaaatttg aagatgaaga acttcatcct aaagcatacg
                                                                       300
ggtcctggca tcttgtccat ggcaaatgct ggacccaaca caaatgggtc ccaatttttc.
                                                                       360
atctgcactg gccaagactg antggttgga tggcaaanca tgtngtgntt ggccaaagtg
                                                                       420
aaagaaggca tgaatattgt ggaaggccat ggaacgcttt tgggtncnag gaatggcaag
                                                                       480
aaccnccagg aagaatcacc cnttnttgac tggggacaac tcnaataagt tgacttgggg
                                                                       540
nttaatntaa ccccccanca attccttttg gaactcagga aacacccttc ancccanttn
                                                                       600
tttcaanttc caaaannttg ggcctn
                                                                       626
      <210> 251
```

. . . . . . . . . . .

```
<211> 603
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (603)
       <223> n = A,T,C or G
       <400> 251
acttttttt tttttttt tttttttc aacagaagaa cttttngttt ctttatttc
                                                                                60
aatattngtc ttattaatat ttttcttatt ttataatgca attacaacaa tttaggagac
                                                                               120
aaaacantat aaacaaaaga atgttaaata gtttttttta aaaaatagct tgttgcttgc
                                                                               180
aagaaagtcc atataatctt attccccccc aaatataatt ttatactttg cactaaacca
                                                                              240
aaatagetta tggaaaatta ggtattaaat agetaaacac agaaaaceta cagetataaa
                                                                              300
taacataaaa tacagtttaa ctttaatgng atgcttaaac aaagcaaact atgatgcant atgaatcaac ttcattaatt ggacaagtcc agtgaggcnc aaattagata agcnctaaac
                                                                               360
                                                                              420
cctcatgatg ggcaagtgaa accttcaccc cagcaagggt ctttcnggtc ttggctatgc caattcettc canaaaagnc ccagttttac angtctggct ttttccgggg gaaccccca
                                                                              480
                                                                              540
tttntttnnc ccaagttggt tnggatttgg cccccannaa atttttttg gngnaaaaan
                                                                              600
                                                                              603
       <210> 252
       <211> 500
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(500)
      \langle 223 \rangle n = A,T,C or G
       <400> 252
aactggaacg gtgaaggtga cagcagtcgg ttggagcgag catccccaa agttcacaat
                                                                              120
gtggccgagg actttgattg cacattgttg tttttttaat agtcattcca aatatgagat gcattgttac aggaagtccc ttgccatcct aaaagccacc ccacttctct ctaaggagaa
                                                                              180
                                                                              240
tggcccagtc ctctcccaag tccacacagg ggaggtgata gcattgcttt cgtgtaaatt
                                                                              300
atgtaatgca aaatttttt aatcttcgcc ttaatacttt tttattttgt tttattttga
                                                                              360
atgatgagee ttegtgeece ecetteece ttttttgtee eceaacttga gatgtatgaa
                                                                              420
ngcttttggt ctccctggga agtgggtgga ngcagccagg gcttacctgt accttggccg
                                                                              480
cgaacaccta aggccaantt
                                                                              500
      <210> 253
      <211> 634
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (634)
```

<223> n = A, T, C or G

```
<400> 253
tegageggee ngecegggea ggtactatta gecatggtea aaccecacce gtgttetteg
                                                                        60
acattgcccg tegacggcga accettgggc ccgcgtctcc tttgagctgt ttgcagacaa
                                                                       120
ggtcccaaag acagcagaaa attttcgtgc tctgagcact ggagagaaag gatttggtta
                                                                       180
taagggttee tgettteaca gaattattee agggtttatg tgteaggggt ggtgaettea
                                                                       240
cacgccataa tggcactggt ggcaagtcca tctatgggga gaaatttgaa gatgagaact
                                                                       300
tcatcctaaa gcatacgggt cctggcatct tgtccatggc aaatgctgga cccaacaca
                                                                       360
atggttccca gtttttcatc tgcactgcca agactgantg gttggatggc aaacatgtgg
                                                                       420
tgtttggcaa antgaaagaa ngcatgaata ttgtggaagc catgganccc tttnggtcca
                                                                       480
ggaatggcag aacnnccagg aanacaccct tgntgactgt ggcaactcga ataaattgac
                                                                       540
ttggggttat cttaacence caacatteet ttggaettag gaancaneee tteaneeent
                                                                       600
tggttcaant tcccaaaaat ttgggctncc tnng
                                                                       634
      <210> 254
      <211> 602
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(602)
      <223> n = A, T, C or G
      <400> 254
60
tggtccttaa gctggaantg caqtgggcac aatcatggnt cactgnagtc tnaacctncc
                                                                       120
aggitcaagi gaicciccia ccicaccicc aniagcigg attacaggca taigcgacca
                                                                       180
tgcccagcta attttttatt ttttgtaaaa acggggtctc actatgtcgc ccangctggn
                                                                       240
cttgaactcc tgaactcaag tgaccettcc gnetnacetn caaagtgeta ggettacagg tgtgaaccac catgectgge ctaaaaaatt tattttaaaa aagtaattta tetettacag
                                                                       300
                                                                       360
ttgtggaggc tgagaaatcc aangncaant ggcncatttg gtgaaaacct tnttgctggt
                                                                       420
ggggactetg tgaaatneec aantggenea tgeatnacae antgangggg ettacattee
                                                                       480
aacatgctat ctcttttaag ttttaaagta cnggccnaaa tntgaacntg aatgacttna
                                                                       540
aatccacnca ttccnctttt ggacnaaaaa ccntgggcaa ttgggatctt ggcnttttna
                                                                       600
aa
                                                                       602
      <210> 255
      <211> 614
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(614)
      <223> n = A,T,C or G
      <400> 255
cgaggtacag gtaagccctg gctgcctcca cccactccca gggagaccaa aagccttcat
                                                                        60
acateteaag ttgggggaca aaaaaggggg aagggggge acgaaggete atcatteaaa
                                                                        120
ataaaacaaa ataaaaaagt attaaggcga agattaaaaa aattttgcat tacataattt
                                                                       180
acacgaaagc aatgctatca cctcccctgt gtggacttgg gagaggactg gaccattctc
                                                                       240
cttagagaga agtggggtgg cttttaggat ggcaagggac ttcctgtaac aatgcatete
                                                                       300
atatttggaa tgactattaa aaaaacaaca atgtgcaatc aaagtcctcg gccacattgt
                                                                       360
```

```
gaactttggg ggatgctcgc tccaacccga ctgctgtcac cttcaccqqt ccaqttttta
                                                                           420
aatcctgagt caaqccaaaa aaaaaaacc anaccaaacn nanaaaccaa ttaagccatg
                                                                           480
ccaatctcat ctggtttctg cncaagtang gttgncaaaa aagggttacc ncactaantc
                                                                           540
ntagccccta aaccnttgcg ggggncantg angggccgan tttganactc cggntggtga
                                                                           600
nccanttggn ggag
                                                                           614
      <210> 256
      <211> 308
      <212> DNA
    . <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (308)
      <223> n = A, T, C \text{ or } G
      <400> 256
nentecagea gtgggteatt egneaacgaa agtentaceg tagaaaagat ggegtgttte
                                                                            60
tttattttga agataatgca ggagtcatag tgaacaataa aggcgagatg aaagggtctg
                                                                           120
ccattacagg accagtagca agggaatgtg cagacttgtg gccccggatt gcatccaatg
                                                                           180
ctggcagcat tgcatgattc tccagtatat ttgtaaaaaa taaaaaaaaa ctaaacccaa
                                                                           240
aaaaaaaaat nnnannnaac annnnanaaa aannnnaaaa aaaaaaagta cctnggccgn
                                                                           300
gaccacgc
                                                                           308
      <210> 257
      <211> 602
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (602)
      <223> n = A, T, C or G
      <400> 257
gcgtggtcgc nggccgaggt acgcggggga gacaaaccat accatatccc accagagagt
                                                                            60
egeagacact atgetgeete catggeeetg eccagtgtat ettggatget gettteetge
                                                                           120
ctcatgctgc tgtctcaggt tcaaggtgaa gaaccccaga gggaactgcc ctctgcacgg
                                                                           180
atcogctgtc ccaaaggctc caaggcctat ggctcccact gctatgcctt gtttttgtca
                                                                           240
ccaaaatcct ggacagatgc agatctggcc tgccagaagc ggccctctgg aaacctggtg
                                                                           300
tetgtgetea ntggggetga gggateette gtgteeteee tggtgaagag cattggtaac agetacteat aegtetggat tgggeteeat gaeeecaaca agggeaeega aeceaatgga
                                                                           360
                                                                           420
aaangntggg antggaataa cantgatgtg atgaattact ttgcatggga gagaaatcct
                                                                           480
tcancatttt naaccccggc cctgtccaac ctntcaaaaa cncacatttt taaggggaaa
                                                                           540
attttactgg atggganggt accetttint ggaagtactg cttttengga nggaagtace
                                                                           600
CC
                                                                           602
      <210> 258
      <211> 600
      <212> DNA
      <213> Homo sapiens
      <220>
```

```
<221> misc feature
      <222> (1) ... (600)
      \langle 223 \rangle n = A,T,C or G
      <400> 258
ggtgtntgng nettatntgt ageggegegg ntggttetga aategeette ageggegeg
                                                                        60
cagtentatt atgtgnatgt cectaceaen aaaatneaga ttaattggna tgeteattae
                                                                        120
ccacgtgaac gccaaagccc ttcgaagtag tgctgccctg cactnaatca agaagttgca
                                                                        180
ttaaaattag aaccaaatcc agagtcactg gaactttctt ttaccatgcc ccanattcag
                                                                        240
gatcagacac ctagtccttc cgatggaaag cactagacaa agttcacctg agcctaatag
                                                                       300
teccagtgaa tattggtttt atggggatag gtgatatggn caatgaatte aagttggaat
                                                                       360
tggnagaaaa actttttgct naagacneng aagenaagaa eccattttet actnaaggea
                                                                        420
cagatttaga cttggagatg gtagcttcct atatccaatg gatgatgctt tcagtccgtn
                                                                        480
cnttgatcag tgncacnttn gaaagcagtt cccaagneet gnaacccagt cctaagccaa
                                                                        540
gtccggttcn gcgattaatc cgactatgta tgcccttcat ngcccctgtn ataaacnggn
                                                                       600
      <210> 259
      <211> 600
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(600)
      <223> n = A, T, C or G
      <400> 259
gccgaggtac atgggaaagg gagtatggng agctatttcc tttttaaagg atgaagacct
                                                                        60
tcataaattg gcccctcgga ttctggtgat tcccgcccgc aagcgcaaat gctccagtgn
                                                                        120
gttatgaaaa tgnttgntaa tctgctctgg ttcttcactg gattcaagan tcgggaggnc
                                                                       180
ttctcgaatc ttttggataa nctggtttaa aacctgaatt gntacccgca tcattttcct
                                                                       240
tttcataaaa atagatatat ctgntcagaa tttctatnaa aagctgcact tgtaganang
                                                                       300
ggtccatgca ctgatttgct atttttaaag cttttttan gcactccatt accctnttgc
                                                                       360
cttcgtgaaa cttcttccca tttttgnccn ggttctggcn gaccngaaga aatgtgccca
                                                                       420
agtgcttaca agttnggcct gacaaggttc nttaaaantt tggatgtacc aagggcccc
                                                                       480
tgggtcctca aaggtcatga atcttttac tggaaccctt atcctttnaa aaggccatgg
                                                                       540
tcaagggaat gnncttcttg gctttgaaac ccggattaan tttttncaaa aaaagccngn
                                                                       600
      <210> 260
      <211> 593
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (593)
      <223> n = A,T,C or G
      <400> 260
acgcgggaac tccatcctca ccacccacac caccctggag cactctgatt gtgccttcat
                                                                        60
ggtagacaat gaggccatct atgacatctg tcgtagaaac ctcgatatcg agcgcccaac
                                                                       120
ctacactaac cttaaccgcc ttattagcca gattgtgtcc tccatcactg cttccctgag
                                                                       180
atttgatgga gccctgaatg ttgacctgac agaattccag accaacctgg tgccctaccc
                                                                       240
```

```
ecgeatecae tteetetgge cacatatgee ectgteatet etgetgagaa ageetaceat
                                                                     300
gaacagetta etgtageaga gateaceaat gettgetttg agecageeaa ecagatggtg
                                                                     360
aaatgtgacc ctcgccatgg taaatacatg gcttgctgcc tggtataccg tggtgacntg
                                                                     420
ggtnccaaag atgtcaatgc tgccttggca ccattcaaac caagcgcaga ttcaatttgg
                                                                     480
ggatggtgcc cactggcttt aaggtngnat naactaccag cttccactgn ggnnctggtg
                                                                     540
gaaactngcc aaggnncctt ggccggaaca ccctangggg aattcanncc act
                                                                     593
      <210> 261
      <211> 343
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(343)
      <223> n = A, T, C or G
      <400> 261
cctacctctc ttnccactgc aaatttctgg gatagaccaa aagtgaattt gattatgtgt
                                                                      60
tggctgaagt tetteattet gaetgttgan gggaggtttt cetttgaaga gtttteatee
                                                                     120
cagactcage tgtcttttca catggatgaa ataatteetg ctaccaacaa cagagettca
                                                                     180
ccaggaagtt gagttttcaa gatgccttgt tgctttgaag aagggagtga tgtcaattct
                                                                     240
cttgntacat tetecettta geaacetgag taagagaete tetgeeactg ggetgeaaaa
                                                                     300
aaataaatta cttgaatctc cccttggccc angctgaggt acc
                                                                     343
      <210> 262
      <211> 593
      <212> DNA
      <213> Homo sapiens
     <220>
     <221> misc feature
      <222> (1) ... (593)
     <223> n = A, T, C or G
     <400> 262
60
ttttttttt ttacagngtn ttttcatttt tattactcaa aaaagtttca ttttttnat
                                                                     120
ttanctttnt gactnigggc tigggccttn aacantttca naacgatttt nigctccicg
                                                                     180
anaaggaaag enecettgat eetgneacna aenentttag eneacatgga aeenecatag
                                                                     240
gecetgntga catgtttett tgtttnggae aatnteataa aaaetttagg nnttacagea
                                                                    300
cnaacceetn naagtnigee tgggeneaca ecanatgeaa attitgggge titteccaace
                                                                    360
ttnttggnat aaaggtaaac aattttatta ccagggggtt cgggacaacc tanttttgtt
                                                                     420
aaaggctgta ttgtaggaaa acctacctcg ggatgtcaaa cccttnacca ttttgagggn
                                                                    480
ctggaaanaa ngttcccgga aanccccggg tancttnggc cggaaccccc taangggnga
                                                                    540
attccnacen ettgggggen gtantaaggg ganceaantt gggecaaant tgg
                                                                    593
     <210> 263
     <211> 591
     <212> DNA
     <213> Homo sapiens
     <220>
```

```
<221> misc feature
      <222> (1) ... (591)
      <223> n = A, T, C or G
      <400> 263
accaagagtt tgctcctggc tgctttgatg tcagtgctgc tactccacct ctgcggcgaa
                                                                         60
tcagaagtaa gcaactttga ctgccgtctt ggatacacag accgtattct tcatcctaaa
                                                                        120
tttattgtgg gcttcacacg gcagctggcc aatgaaggct gtgacatcaa tgctatcatc
                                                                        180
tttcacaaag aaaaagttgt ctgtgtgcgc aaatccaaaa cagacttggg tgaaatatat
                                                                        240
tgtgcgtctc ctcagtaaaa aagtcaagaa catgtaaaaa ctgtggcttt tctggaatgg
                                                                        300
aattggacat agcccaagaa cagaaagaac cttgctgggg ttggaggttt cacttgcaca
                                                                        360
tcatggaggg tttaatgctt atctaatttg tgcctcactg gacttgncaa ttaatgaagt
                                                                        420
gatcatattg catcataagt ttgctttggt taancttaca ttaaagttaa ctggatttta
                                                                        480
agggaattat actgtaggtt ctggggtaac tatttaatac taattttcat aacnattttg
                                                                        540
gttaatncca agttnaaatt tatttggggg gaanaaaatt tttggccttc t
                                                                        591
      <210> 264
      <211> 595
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (595)
      <223> n = A, T, C or G
      <400> 264
accaagagtt tgctcctggc tgctttgatg tcagtgctgc tactccacct ctgcggcgaa
                                                                        60
tcagaagtaa gcaactttga ctgccgtctt ggatacacag accgtattct tcatcctaaa
                                                                        120
tttattgtgg getteacaeg geagetggee aatgaagget gtgacateaa tgetateate
                                                                        180
tttcacaaag aaaaagttgt ctgtgtgcgc aaatccaaaa cagacttggg tgaaatatat
                                                                        240
tgtgcgtctc ctcagtaaaa aagtcaagaa catgtaaaaa ctgtggcttt tctggaatgg
                                                                        300
aattggacat agcccaagaa cagaaagaac cttgctgggg ttggaggttt cacttgcaca
                                                                        360
tcatggaggg gtttagtgct tatctaattt gtgcctcact ggacttgtcc aattaatgaa
                                                                        420
gttgattcat attgcatcat agtttgcttt ggttaagcat cacattaaag ttaaactgga
                                                                        480
ttttatggta tttatagctg nanggtttct ggggttanct atttaatact aaatttccat
                                                                        540
aagetttttg ggttaangee aagnttaaaa tttttttggg ggggaaaaaa atttt
                                                                        595
      <210> 265
      <211> 592
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (592)
      <223> n = A, T, C or G
      <400> 265
ggtacttttt ttttttttt tttttttt ttgaaaatta tacttttatt tgagtcacca
                                                                        60
ggagaaagat teacttgtgg tteaagteaa atgtteanaa teataacagg eeanaaaggt
                                                                        120
ttgatecega geacaageee aegagggagg ggaccaaaae agaccaaaat gagacaacaa
                                                                        180
ccccatataa aaagatgaac tggcggcttc acacactcac acacatacac atacacacgg
                                                                        240
```

```
atgaaatgtt tggacagagg caaatttcac gtggtcattt ctgtttcttt ttaaatacag
                                                                             300
gtttgtgggg tggtattttg tttttccag ctataaaaaa aggcccaaaa gtgcatgtgt
                                                                             360.
gagggggaa aggcagaaat taagcaataa agtcattttc cctggaggga catganaggg
                                                                             420
agaaaacagg aggcaattgc tggganaacg cactttctta acactqqqct tttqqqtatt
                                                                             480
cttantattg gnccncaaaa agttattttc acattctaac tttgaagnet ntttccnggg
                                                                             540
attnaatggn ccttaaaacc tttgggaact ttaaaaaaac cngggcttac cc
                                                                             592
       <210> 266
       <211> 594
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(594)
       <223> n = A, T, C \text{ or } G
       <400> 266
acgcggggaa aaaaaaggca gtattccctt tttaaatgag ctttcaggaa gttgctgaga
                                                                              60
aatggggtgg aatagggaac tgtaatggcc actgaagcac gtgagagacc ctcgcaaaat
                                                                             120
gatgtgaaag gaccagtttc ttgaagtcca gtgtttccac ggctggatac ctgtqtqtct
                                                                             180
ccataaaagt cctgtcacca aggacgttaa aggcatttta ttccagcgtc ttctagagag
                                                                             240
cttagtgtat acagatgagg gtgtcccgct gctgctttcc ttcggaatcc agtgcttcca
                                                                             300
cagagattag cotgtagott atatttgaca ttottcactg totgttgttt acctaccgta
                                                                             360
getttttace gttcacttcc cettccaact atgtcccaga tgtgcagget cetectet
                                                                             420
ggactttctn caaaggcact tgaccettcg gnetetaett ggeceetnae eteaceeet
                                                                            480
tetggeaceg gnentgngae atteactten gagaagaeen eeceeaagga ggenggegnt
                                                                            540
tggnccanga aaaaaccccg gggaagggtt tnttttttn aaagggaaat ttcc
                                                                            594
       <210> 267
       <211> 598
      <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (598)
      <223> n = A, T, C or G
      <400> 267
actggccctc ggtgctggca aaggtgtagt tccactggcc gagggaatca agacatagtg
                                                                              60
gtccttctgc taagccaagg gctgccacaa tgacacagta gccagatcct gcaattccaa
                                                                             120
tgagagcagc caatacagaa gaaagcatcg cacatcgttt gccacagttt tcatggccacagcagcacaagcacaagcatca tgttccagcc caatgaagac aaatgctggc aggagcatca
                                                                            180
                                                                            240
gcagggccac ctcctacgat gccagaaaag aaccacacga aacggctgag gtggttttcg
                                                                            300
gaggcatact ttgttcccat tgggaaagta aagccaaata ttacccgcga tgcacaggaa
                                                                            360
ggggcgagcc caaccagaaa atgtccgaat gcatcgtgca cacttcccat agcacatggt ggtcttgcta ggtttttctc ccccttctct ttggncttca acttcagtga taccccaaat
                                                                            420
                                                                            480
tagatgaaag tggtgccctt ttgggtggaa aaagcaaaca ccaaccccgg gtacctttgg
                                                                            540
geoggaacae nettaaggee aatteeanne aattggegge eegtaettan gggateee
                                                                            598
      <210> 268
      <211> 590
```

```
<212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (590)
      <223> n = A, T, C \text{ or } G
      <400> 268
ggacatatta tcaataggct ataagatgta acaacgaaat gatgacatct ggagaagaaa
                                                                         60
catcttttcc ttataaaaat gtgttttcaa gctgttgttt taagaagcaa aagatagttc
                                                                        120
tgcaaattca aagatacagt atcccttcaa aacaaatagg agttcaggga agagaaacat
                                                                        180
ccttcaaagg acagtgttgt tttgaccggg agatctagag agtgctcaga attagggcct
                                                                        240
ggcatttgga atcacaggat ttatcatcac agaaacaact gttttaagat tagttccatc
                                                                        300
actotoatoo tgtattttta taagaaacao aagagtgoat accagaattg aatataccat
                                                                        360
atgggattgg agaaagacaa atgtggaaga aatcatagag ctggagacta cttttgtgct
                                                                        420
ttacaaaact gtgaaggatt gtggtcacct ggaacaggtc tncaatctat gtagcactat
                                                                        480
gtggctcanc cttggtaccc cttggattat atatcaacct gnaacatgng nctgggactt
                                                                        540
actitionaaa chaaathtto otthtitigaa gaaaatotgg gtttttghaa
                                                                        590
      <210> 269
      <211> 602
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(602)
      <223> n = A,T,C or G
      <400> 269
acttgaagga agtcgaatca gagatagact ctgaagaaga acttataaat aaaaaaagaa
                                                                         60
tcatagagaa agttattcat cgactcacac actatgatca tgttctaatt gagctcaccc
                                                                        120
aggetggatt gaaaggetee acagagggaa gtgagageta tgaagaagat eectaettgg
                                                                        180
tagttaaccc taactacttg ctcgaagatt gagatagtaa aagtaactga ccagagctga
                                                                        240
ggaactgtgg cacagcacct cgtggcctgg agcctggctg gagctctgct agggacagaa
                                                                        300
gtgtttctgg aagtgatgct tcaggatttg ttttcagaaa caagaattga gttgatggtc
                                                                        360
ctatgtgtca cattcatcac aggtttcata ccaacacagg cttcagcact tncntttggt
                                                                        420
ggtggttcct ggtcccntgg aagttggaac caaattaatg gngtagtctc tatacccaat
                                                                        480
acctttggtt ttcatgtgta anaaaaaggn ccattacttt taanggattg tgctggnctt
                                                                        540
attgngccan taactttttt ttaaatggcc cagttacngg ttttaattct taaaannaaa
                                                                        600
                                                                        602
      <210> 270
      <211> 595
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(595)
      <223> n = A,T,C \text{ or } G
```

```
<400> 270
ggtacgcggg ggtaggagcc teteteceta etgetgetae acaagaccet gagactgace
                                                                            60
tgcaggacga aaccatgaag agcetgatee ttettgecat eetggeegee ttageggtag
                                                                           120
taactttgtg ttatgaatca catgaaagca tggaatctta tgaacttaat cccttcatta
                                                                           180
acaggagaaa tgcaaatacc ttcatatccc ctcagcagag atggagagct aaagtccaag
                                                                           240
agaggateeg agaacgetet aageetgtee acgageteaa tagggaagee tgtgatgaet
                                                                           300
acagactttg cgaacgctac gccatggttt atggatacaa tgctgcctat aatcgctact
                                                                           360
teaggaageg cegagggaee aaatgagaet gagggaagaa aaaaaatete tttntttetg
                                                                           420
gaggetggca cetgattttg tatececetg tageageatt actgaaatac ataggettat
                                                                           480
atacaatget tetttetgga tattetettg gettgggtgg acceetttt ceggeceag aattgttaan taatngaann neentneann aagggnnnaa aggnaaatea nettt
                                                                           540
                                                                           595
       <210> 271
       <211> 592
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(592)
      \langle 223 \rangle n = A,T,C or G
      <400> 271
ggtacattga gatecegeet etacaaaate aaaaaattag eeaggeaagg tggtgegtge
                                                                           60
ctgtcgcccc agctacttgg caggctgagc tcaggaggtc aagcctgcct tgggccatga
                                                                          120
teateceatg caetecagee tgacatteag ageaagaeet tgteteaaag aaagaaaaae
                                                                          180
atttttatgg tgttttcttt tttagtcttt tcaataatga aaattttcat tttacaggta
                                                                          240
aaatgaaagg cctggcattt attcaagatc ctgatggcta ctggattgaa attttgaatc
                                                                          300
ctaacaaaat ggcaacctta atgtagtgct gtgagaattc tcctttgaga tttcagaaga
                                                                          360
aaggaaacaa tgtgattcaa gatatttaca taccagaagc atctaggact gatggatcac
                                                                          420
tgtcccgatt caaattattc ttcagtccat ttcccctttc tatttcagct ggtccttttc
                                                                          480
acctaactgt cagtcattct ggtttcaacn atgctttatc tcatgtcctt gaatatagtt
                                                                          540
ggggnacttt aatttttang gaataatnna acagnttccn ttaaaggntn ng
                                                                          592
      <210> 272
      <211> 607
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(607)
      <223> n = A,T,C \text{ or } G
      <400> 272
acattaaagt gtgatacttg gttttgaaaa cattcaaaca gtctctgtgg aaatctgaga
                                                                           60
gaaattggcg gagagctgcc gtggtgcatt cctcctgtag tgcttcaagc taatgcttca
                                                                          120
teetetetaa taaettttga tagacagggg etagtegeac agacetetgg gaageeetgg
                                                                          180
aaaacgetga tgettgtttg aagateteaa gegeagagte tgeaagttea teecetettt
                                                                          240
cctgaggtct gttggctgga ggctgcagaa cattggtgat gacatggacc acgccatttg
                                                                          300
tggccatgat gtcaggctcg gcaacaggct ccttggtgac actcaccaca ttgnttttca
                                                                          360
agetgaettt cagettgnca cettggagag aetttaacce ggaecaaggg eeegatgeet
                                                                          420
teegttacee aggaatttea teaceaatgg tggtanttea ggaatgttgg caagttteet
                                                                          480
```

```
tggcatnttc ccaaanagtt tgttcccgtt cttnttgggn ggcangggct tcggaaaggg
                                                                         540
ttnattttgt ngggaacena aaaactgggg tnaaacteet tneeggttna ngggttteeg
                                                                         600
nnanccn
                                                                         607
      <210> 273
      <211> 398
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(398)
      <223> n = A,T,C or G
ggtaccgcca ttattctttt gggcaccttt ggttgttttg ctacctgccg agcttctgca
                                                                          60
tggatgctaa aactgtatgc aatgtttctg actctcgttt ttttggtcga actggtcgct
                                                                         120
gccatcgtag gatttgtttt cagacatgag attaagaaca gctttaagaa taattatgag
                                                                         180
aaggetttga ageagtataa etetacagga gattatagaa geeatgeagt agacaagate
                                                                         240
caaaatacgt tgcattgttg tggtgtcacc gattatagag attggacaga tactaattat
                                                                         300
tactcagaaa aaggatttcc taagagttgc tgtaaacttg aagattgtac ctqccccqqq
                                                                         360
ccgnccgctc gaaagcttaa ntggccgttt cnaanncg
                                                                         398
      <210> 274
      <211> 587
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (587)
      <223> n = A, T, C or G
      <400> 274
acttttttt tttttttt tttgttgaat caaaagcagg gtttattttt ctatcaaatc
                                                                          60
cccaatccat gttccagcca atggatgaag ggtgaatcaa gccccacata gactcttggt
                                                                         120
aaaaaaaaaa aaagccaaca cactttttc tttctttca
                                                                         180
aaaagctccc aggcctttgg gaacagctga aacaaattca tatcctgact aggtctgttt tctcttaggt atttggatgg tccctctctg ctgccacttc tgcacagatg aggcactgat
                                                                         240
                                                                         300
aatggcetge aggtcactca caatcetage tecacatcae tecatggttt gataacetag
                                                                         360
aaccacgtta tgatttccat ttataatgcc ctaagaacag ctgaaaagat ctgtattaaa
                                                                         420
ttctgcaaat ctttattgag tgccactatt tgctgggcac angctaggen ctggattctg
                                                                         480
ctggttcttg agaaacctaa aanggnncct tnggccggaa caccettang gcgaaatcca
                                                                         540
cncactgggg ggcgtactaa ngggatccaa ctttggncca acttggg
                                                                         587
      <210> 275
      <211> 588
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (588)
```

... ... ... ...

360

```
<223> n = A, T, C or G
       <400> 275
 acttttttt tttttttt tttgccttta taagagaatt tttattgtta attatttacc
                                                                           60
ttaatagttt cagaaagagg aacaaattag ctcagtccaa catgattggc agttggcata ttctagtgaa gcaagtgttc tgactgctaa ggatttaatt tggataattt taatacttag
                                                                          120
                                                                          180
 ccatctaaca cttcaagcat aacccagaat aaatgcacca ccttcctttc actttaatac
                                                                          240
ccgnacctac ctcacttcga tataagaaat atcattcaat atgatttcca gaagggacaa
                                                                          300
gtttcctgga gaatacaggc atganggaca atgcacaaaa agaaaaactc aaaatnaaac
                                                                          360
 tctggatgga taattactaa gctaagggaa ccaaaccttc caatttntaa agaaattaaa
                                                                          420
teeggtteea aatgeetnat angnetatgt tnaaaaggtt etggattaat aceggaaaag
                                                                          480
gnttgnttnt tacaggatnc cccaaccgtt acgggccctt ngcccagaat gggccttaaa
                                                                          540
anccaaagng tetttteegn ngaggeecea tttnanaate ettntttt
                                                                          588
       <210> 276
       <211> 595
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
       <222> (1)...(595)
       <223> n = A,T,C or G
      <400> 276
actttagata catcattcct caaaaagttt ttaacggaga aagtggggca attcaatggg
                                                                           60
ggaaaggacg gcetttttaa caaatggtgc tggttctact gggtatctgc atcettgata
                                                                          120
cacagaagtt aactcaagat ggaccacaga ctcacatgta agagctaaaa taacattett
                                                                          180
agaagaaatc atggaagtaa atcttcgtga ccttggatca ggtaatgggt acttttttt
                                                                          240
tittitttt ttittta tcagattaat tttacittat ticttcaggc ctggggtttt
                                                                          300
tcgatgactt caaatttggg atcttcaaat ttgaaggtgg gaaatggtat tcatgtctgc
                                                                          360
attaccaaac atttgctttg acttaaaaag ctcctctcca gctcttgccg atctctgaac
                                                                          420
tagcatcaac aggnteetee agatgtetgg neettaaatt tggatteeet aatettggee
                                                                          480
acaaagangt ttettggata gggaacaaag tteeettatt naaatgeean tngtngaace
                                                                          540
nccaatgttc cttcncaaaa ngggcttaaa ccggttaccc aattgacaaa ggaaa
                                                                          595
      <210> 277
      <211> 597
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(597)
      <223> n = A, T, C \text{ or } G
      <400> 277
ggtactgttc ctgttggccg agtggagact ggtgttctca aacccggtat ggtggtcacc
                                                                          60
titgetecag teaacgttac aacggaagta aaatetgteg aaatgcacca tgaagetttg
                                                                         120
agtgaagete tteetgggga caatgtggge tteaatgtea agaatgtgte tgteaaggat
                                                                         180
gttcgtcgtg gcaacgttgc tggtgacagc aaaaatgacc caccaatgga agcagctggc
                                                                         240
ttcactgctc aggtgattat cctgaaccat ccaggccaaa taagcgccgg ctatgccct
                                                                         300
gtattggatt gccacacggc tcacattgca tgcaagtttg ctgagctgaa ggaaaagatt
```

```
gatcgccgtt ctggtaaaaa gcttggaaga tggccctaaa ttcttgaagt ctggtgatgc
                                                                       420
tgccattggt tgatatggtt cctggcaagc ccatgtgtgt tgaaagcttc ttaaactatc
                                                                       480
cacctttggg tcgctttgct ggtccngatt tgagacanac catttccgnn gggtggcaat
                                                                       540
caaaccattg ggccaanaaa gnttntggac ttgcaagggn nccaaatttt ncccaaa
                                                                       597
      <210> 278
      <211> 595
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(595)
      <223> n = A, T, C \text{ or } G
      <400> 278
ggtacttttt tttttttt tttttttt ttagtttatt aaaatactga gttttatttc
                                                                        60
acatgtatat ttttgtctcc ccaccatttc catgtctgac caccgctact actatgtcct
                                                                       120
atcataacat tccatacata cttaaaacca agcaaagggt ggagttccat ctttaaaaac
                                                                       180
taaacaggca ttttggacaa cacattcttg gcaatagaac ctggacaaca tttatcaaac
                                                                       240
acggtaggga aagttctcac tctgcattat aaaaaggaca gccagatatc aactgttaca
                                                                       300
gaaatgaaat aagacggaaa attttttaac aaattgntta aactattttc ttaaagagac
                                                                       360
tteeteeact gecagagate ttgaatagee tettggneag teatteegga aacaattett
                                                                       420
ccataattga tgaatttggc tttcactttt gggaagagaa cccccttttc tatacttggg
                                                                       480
tgcattttgc ttaaaggctt ctacaaacta gggcctttgg gggtttaaga gttttccngg
                                                                       540
gtcttgaagg ntcttggcct ttgaacttgg ggtnaaaang gttgngcttt tccat
                                                                       595
      <210> 279
      <211> 586
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(586)
      <223> n = A,T,C or G
      <400> 279
ggtacgcggg gagatacgtt cgtcagcttg ctcctttctg cccgtggacg ccgccgaaga
                                                                        60
agcatcgtta aagtctctct tcaccctgcc gtcatgtcta agtcagagtc tcctaaagag
                                                                       120
cccgaacage tgaggaagee etteattgga gggttgaget ttgaaacaae tgatgagage
                                                                       180
ctgaggagcc attttgagca atggggaacg ctcacggact gtgtggtaat gagagatcca
                                                                       240
aacaccaagc gctccagggg ctttgggttt gtcacatatg ccactgtgga ggaggtggat
                                                                       300
gcggctatga atgcaaggcc acacaaggtg gatggaagaa ttgtggaacc aaagagagct
                                                                       360
gtctccagag aagattctca aagaccaggt gcccacttaa ctgtgaaaaa agatatttgg
                                                                       420
tggtggcatt naagaagacc ttgaagaaca tcacctaaga gattattttg acagtatgga
                                                                       480
aaattgaatg attgaaatca tgacttgacc aagcatggcc aaaaaagggc tttgctttga
                                                                       540
accttgagac atgattengg ataaaatgen tenaatnent ntggga
                                                                       586
      <210> 280
      <211> 612
      <212> DNA
      <213> Homo sapiens
```

```
<220>
       <221> misc feature
       <222> (1)...(612)
       <223> n = A, T, C \text{ or } G
       <400> 280
60
gtcatgaagg ccatggggtt ggcttgaaac cagctttggg aggttcgatt ccttcctttt
                                                                           120
ttgtctaaat tttatgtata cgggttcttc aaatgtgtgg tagggtgggg ggcatccata
                                                                           180
tagccactcc aggtttatgg agggttcttc tactattagg acttttcgct tnaaaacgaa ggcttntcaa atcatgaaaa ttattaatat tactgctgtt anaaaaatga atgagcctac
                                                                           240
                                                                           300
anatgatagg atgtttcatg gggngtatgc atcggggtaa tccnaataac gtcggggcat
                                                                           360
tccggatagg cccaaaaang tttntgggaa aaaaagtttn atttaccccc attaaattta
                                                                           420
tnnnnaaaag ggattttgcc taaggttggg ctaagggggt anccengaaa attgggggaa
                                                                           480
atcangnaat gaaaccccct ntgatggnca aaaacagctc ctnttggttg ggccttatng
                                                                           540
ggaanngggc ttcaacntan naccttnggc ggnaaaaccc ttanggngaa ttnnnnncaa
                                                                           600
ntggggggg tn
                                                                           612
      <210> 281
      <211> 593
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (593)
      <223> n = A, T, C or G
      <400> 281
acgctgcttc ttcagagcaa tacgccgccg tttgtgctgc aggacacgtg gagtaacaag
                                                                           60
acgetgaate ttgggtgett tggteetagg tttettaeet tetttattta agggetttet
                                                                           120
tacaacatac tggcggacat catettettt agagagattg aaaagtttgc ggattetget
                                                                           180
agetettttg gggeecagge ggegaggeae tgtagtatea gteagteeag gaataceett
                                                                           240
ctctcctttt tttacaataa ccaagttgag aacgctcaga tttgcatcca caatgcaacc
                                                                           300
acgaactgat tttctctttc tttctcagtt ctccttggtc tgtaacagga atgcccctta
                                                                          360
ctcaatanca ggcggacacg ggcatgggtc aagacaccct gcttcatggg gaaaccttgg
ttgncgttcc accactggat tcggaccaca taaacctttc attcttnaac caaacgtaac
                                                                          420
                                                                          480
ancaactttt ggnggccata encttttata naaagteegg ggganaagtn ttttgcagga
                                                                          540
caageetgta acnaatagtn aaateeegga tttggattee taaneetttt een
                                                                          593
      <210> 282
      <211> 595
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (595)
      <223> n = A, T, C or G
      <400> 282
ggtacaattc aagaaactaa gtatttatgg gcattgaaga aaaaatgttg agataaaatt
                                                                           60
```

\*\*\*\*\* \*\*\*\* \* \*\* \*\*\*\* \*\*\*\* \*\*\*\* \*\*\*\*

```
gctgtgcaga aaaaagtgtt aatgaagccg acctgactac ttaaccttag agacctgctt
                                                                     120
tacaaggttg gcccttgatt ggcatctggg aacttggagt tcagggggct tccaccattc
                                                                     180
ccagaactga tcaaagtagc ttactatatc taaactgtaa aacaatatag tttctcctga
                                                                     240
acacctgctt tccttctggg agtctggaat tttggtatgt gccaggcaga gactaccttt
                                                                     300
gtgaccaget cccagtaaaa accccaggca ctcagtctct aacaagcttt tctggttgac
                                                                     360
agtgtttcac aagtgctggt acaactggtt gctgggagaa ttaagctcat cctctgtgat
                                                                     420
tecactggcc gaggattett ggaagettgc acttaagttt ceeetgactt caceccatgg
                                                                     480
gettttttee ttgetgattt ggtttgnate etteetgnat aaateatgge etgaacenaa
                                                                     540
cttgaaaaaa aaannnnnn nnaaaaaaag gtncttgccc ggcggccgtt naaat
                                                                     595
      <210> 283
      <211> 348
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (348)
      \langle 223 \rangle n = A,T,C or G
      <400> 283
60
ggtanagggg gtgctatagg gtaaatacgg gccctatttc aaagattttt aggggaatta
                                                                     120
attntaggac gatgggcatg aaactgtggt ttgctccaca natttcanag cattgaccgt
                                                                     180
240
ctgtttttaa gcctaatgtg gggacagctc atgagtgcaa nacgtnttgt gatgtaatta ttatacgaat gggggcttna atcgggagta cctnggccgn naccacnc
                                                                     300
                                                                     348
      <210> 284
      <211> 563
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (563)
      <223> n = A, T, C \text{ or } G
      <400> 284
ggtacccatt aatttgctca gatatagcag gcttaatggt tctatatttt caaaagtttt
                                                                      60
taagaatggt ttctaacgta ggagagggaa aacatccacc atcccttttc agaatttaaa
                                                                     120
tggagggcag taaacattct ttacacccaa aacctatggc agcagttcaa atttgaccaa
                                                                     180
ggtaaatgta gaatagagat gttctaaaca cagctaggac tcagcaagtc taacacacta
                                                                     240
aaatcatatg attacatttt aaaagaaaat gcacaaaaac caaatagaaa ttttgagatt
                                                                     300
ttttttcatt tgaaggtaat cttaatgcta ttaaattcac aaatgctaat ttaaataccc
                                                                     360
aatcctattt atctaaaaca cacattgcaa acacacaaat tatctattct ctccacatgt
                                                                     420
cagccgccca ttcatatcat ggtttggaaa tgggggagaa atagattncc cttaaactgc
                                                                     480
aagtcaacan ggggttettt acagttaact ttagecaaat teataceaaa taeeegggta
                                                                     540
ectgecengg eggeegtten aaa
                                                                     563
      <210> 285
      <211> 422
      <212> DNA
```

```
<213> Homo sapiens
       <400> 285
acaatggact ggatactaga aattttcttt tcactcaaca gaacataggc atcctggaat
                                                                           60
tcacatttct gaccttttga tgtattaata aagtatggag aaatatagcc tcgatcaaac
                                                                          120
tteatgeett caataattte taatteatea tteagtgitt tteeateett taetgtgatg
                                                                          180
acaccettte trecaactt trecattgea teagagatga tattgecaat trettretet
                                                                          240
ccgtttgcag aaatcgtagc aacctgtgca atttcttcag gggtggtcac aggtttagac tgctttttaa gttcagcaat tacagcatca acagctaaca tcacacctct cctgatttcc
                                                                          300
                                                                          360
actggattag cacctttgct aatcttctcg aagccttctt ggctatagag cgtgccagta
                                                                          420
                                                                          422
      <210> 286
      <211> 588
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(588)
      <223> n = A, T, C or G
      <400> 286
actgttcctg caggttaagg caggactgga actcctccac agcttgcaca tagttttcag
                                                                           60
atteaacact aactteteeg agtitaagat gtgeetggge agcataaage tgtgettett
                                                                          120
ttgtttettg cettttaaaa atgatetttg etaaateeag catateeeag geaageteta
                                                                          180
ggttcccaat ctcctcctcc tcattttctt gaagagactt gttttcaagg actgaatcat
                                                                          240
ttggcatttc ttcggtctta tcattttctt tatcatcctc ttctgagcct tcagtttcat
                                                                          300
ctatgttatc attatttct accagagatt catcttctgn tnttttctcc ttcttcctct
                                                                          360
tncacatgca caccttccaa ggcgtttcca acacaccatt cttcatcttg ccaacttcag
                                                                          420
aagtggattt ccatagaaaa agaangnttn ttcacactta ttaactgctc ttcatacttt
                                                                          480
ttacctnaaa gactaactgn ttcctggaat gcattggccg ctgctnggaa atccccatan
                                                                          540
engaagttnt ggeetaance aaagttntta gttactttee cateegae
                                                                          588
      <210> 287
      <211> 583
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(583)
      <223> n = A, T, C or G
      <400> 287
actggaactc caggaagcgc tggccagcct catacgggag ccatttttct ttcactgcct
                                                                           60
ctgctgctga catcttcttc tttcccttca caccctcgaa gcctatgaag gctttctgag
                                                                          120
caggetteag cetggtggee atgtettggt caateacace etgggagaet gegteetgaa
                                                                          180
gtgacagett etggecegtg gttgggtgga tgatgecace tgtgcaggee tgageeteca
                                                                          240
gaageetetg acceptgatg etgteaacga tgeecegete tatacettet gtaatggaga
                                                                          300
ttttctccag gttttctgtg tcaaagatgg ctgcaatggg gctcgattct tncagggtgt
                                                                          360
ctgaaaaaga actgctcctt atggntaaat tcctgacctg gatatggtgg aaatcttact
                                                                          420
tactgattca tgtcgggagc tgctaaaaac atnatcgttg caccactggc catgctgtgn
                                                                          480
```

```
ttggngccac accatttttn angngacatg taacnaattg antaggttag nttccqaacg
                                                                          540
gaccttggcc ggaacaccta aggngatcan ncatggggcg tnn
                                                                          583
      <210> 288
      <211> 607
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (607)
      <223> n = A, T, C or G
      <400> 288
ggtacttttt ttttttttt ttttttgtt atttagttt tatttcataa tcataaactt
                                                                           60
aactetgeaa teeagetagg catgggaggg aacaaggaaa acatggaace caaagggaac
                                                                          120
tgcagcgaga gcacaaagat tctaggatac tgcgagcaaa tggggtggag gggtgctctc
                                                                          180
ctgagctaca gaaggaatga tctggtggtt aagataaaac acaagtcaaa cttattcgag
                                                                          240
ttgtccacag tcagcaatgg tgatcttctt gctggtcttg ccattcctgg acccaaagcg
                                                                          300
ctccatggcc ttcacaatat tcatgccttc tttcactttg ccaaacacca catgcttgcc
                                                                          360
atccaaccac tcagtcttgg cagtgcanat gaaaaactgg gaaccatttg gggttgggtc cagcattttg catggaccan aatgccagga cccctatgct ttaaggatga anntcttatn
                                                                          420
                                                                          480
ttnaaatttc ttcccataaa nggcttgcca ccaangccat tatngcgngt gaagcaccac
                                                                          540
etgacecata accetggaat aattninga aaaceggace ettniacena atcittitte
                                                                          600
aggggnn
                                                                          607
      <210> 289
      <211> 591
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (591)
      <223> n = A, T, C or G
      <400> 289
actititit tittitit titgagaatg aataagcagt totttaatgg ttatttaaat
                                                                           60
atattocaga agagogitta taattoatti acaagigoag tattgogota giaaatqita
                                                                          120
cttgacctct tgtataaata atgccgatta agaattagtc ctggaatagt tttcgaattt
                                                                          180
ctaactctgt agatctaaaa cacaattgta aatggtataa agatgtaaga atcatattgt
                                                                          240
gataaagtca atctcaaaaa tagagaatcc agacccttcc cagataattt aagaactgag
                                                                          300
ttttcctcaa cttaaacatg atggccacac agaaaacagt aaagacactt ttcqatqtqa
                                                                          360
tacaactgga taaaactcga gaatatgagt atttagngac caatgnatan acattantgg
                                                                          420
aattttaaaa neeetttaa tetgaageeg aaaaaaange eatttteeaa gaattattgn
                                                                          480
geoctaatea teatenanne nngaatanna thentteeen ggatagnnnn nnnteeneet
                                                                          540
tnggaaantg ggccnaantt ntttggtntn aaagggggnc cnttaantcc n
                                                                          591
      <210> 290
      <211> 592
      <212> DNA
      <213> Homo sapiens
```

```
<220>
       <221> misc feature
       <222> (1) ... (592)
       <223> n = A, T, C \text{ or } G
       <400> 290
ggacttggaa atggttgtct ggaaagcttc cactttggtc ttgacggcat tcaccctctc
                                                                                  60
cagcacette teetggattg ctaceceaaa ateattteea tetteaatet tggggateag
                                                                                 120
gtgttggatc catgtaatca ccagaatgca tttctctttg agagtccaga cttctggctt
                                                                                 180
aaccagggca agcagggaca ggactttctc attcccaggg agaaatccac acttagggac
                                                                                 240
ttetttette teetgettat etgttteeat eteateatee ttgggtggag ggtetgggat
                                                                                 300
ggggatgtcc agtggggccc ggagggaagt caagtcagcc acattgaggg agtcctcttg
                                                                                 360
caagagetga tteaggtata tgattttetg tggcaagaat etgtagagga attecteane etnetggaaa agaatetgte tgaagacett caeetggttg egggetttee egetaagege
                                                                                 420
                                                                                 480
accccacacg gtttgggcct gctgntttaa tccttaanct ctggcttccg gntagtcccc
                                                                                 540
cgggaccttg ccggccggcc ntcaaagggc aattcancna ctggcggccg tn
                                                                                 592
       <210> 291
       <211> 609
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(609)
       <223> n = A, T, C or G
acagtggcat gatctcggct cactgcaacc tctgcctccc gggttcaagc aattctcctg
                                                                                  60
cetcagecae ecaagtaget gggactaeag gtgegtgeea ecaegeceag etaaattttg
                                                                                 120
tatttttagt ggagacgggg tttcaccatg ttggccagga tggtctcaat ctcctgaccc
                                                                                 180
tgcgatctgc ccacctcagc ctcccaaagt gctgggatta caggcgtaag ccaccgggcc tggcctgttt tatgattctt aatagttact tggtttaaat cacatttgat actatccttc
                                                                                 240
                                                                                 300
tgaaaagtct gagacagatc tacaaactac agtcaaaatt atagattaag aggaatgaat gcacctattt ggctttaagt tgaagatgaa ttatttctca tgctcatttt cttgcngcag
                                                                                 360
                                                                                 420
ttatcttaga aagaccccca aaggcttgtg attgtaaagc acttgcatga tcacagaatg
                                                                                 480
caagettetg gtacettegg cegtgacaeg etaagggega atteateaea attgegggee
                                                                                 540
gtacctatgg atccannete ggteeaactt ggeggaatea tgggeataet gntteetggn
                                                                                600
nnaaatgtn
                                                                                 609
      <210> 292
       <211> 568
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(568)
      <223> n = A,T,C or G
      <400> 292
actgcccaga aggagttcat aaagaataca aagaagaccc caaaagatgt cacgatggca
                                                                                  60
ctattgaatt cacgagcatc gatgcacaca atggtgtggc cccatcaaga cgtggtgatt
                                                                                120
```

```
180
tggaaatact tggttattgc atgatccaat ggcttactgg ccatcttcct tgggaggata
                                                                         240
atttqaaaqa tootaaatat gttagagatt ocaaaattag atacagagaa aatattgcaa
gtttgatgga caaatgtttt cctgagaaaa acaaaccagg tgaaattgcc aaatacatgg
                                                                         300
aaacagtgaa attactagac tacactgaaa aacctcttta tgaaaattta cgtgacattc
                                                                         360
                                                                         420
ttttgcaagg actaaaaact ataggaagta agggtgatgg caaaatggac ctcaatggtg
tggaaaatgg angnttgaaa gccaaaacca tnnnnnaaaa ncttagggcg aattccannc
                                                                         480
actggcggcc gtnctaangg atccagcttg gncccaactt ggggtaatca tgggcataac tggtncctgg ggaaaatggt ttcccnnn
                                                                         540
                                                                         568
      <210> 293
      <211> 603
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (603)
      <223> n = A, T, C \text{ or } G
      <400> 293
60
tittaanaaa cttttatttg agnggntntt acaaanattg nngcaatatg aaagtcattt
                                                                         120
                                                                         180
gtttgatana aatatcaago tgnottgtca aacacnotga agtaacccaa aaatninttt
caaageteae anagettaaa aagagenaag attntntgea accagacaaa acctatttnt
                                                                         240
                                                                         300
gcatttccta tttctttctn aaactgnttt gcctaccaaa ctttnacgtt taaacatttt
caggaaatgc agggatcatt ttgtttggaa ttttaagacc ccccngaacn cataggtntt
                                                                         360
tacaaagaaa cttttcccga tcccttaatt gaaaagaacc ntccnaaata taaantttgn
                                                                         420
aaactcccnt ttttggccaa ttgatcanaa tgccagaaga natgctaacc naanagccct
                                                                         480
                                                                         540
ttaactgggc tgggattcca taccctaaan ggggtttcaa aactggttaa ccttnnccca
                                                                         600
attttaacct tngggaaaag ggnaaaggan ccccggggna aaaataaggt tttgaaaaat
                                                                         603
aaa
      <210> 294
       <211> 617
       <212> DNA
      <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (617)
       <223> n = A, T, C \text{ or } G
       <400> 294
ggtacgcggg gatcgcttcc tggtcctcgc cccctccgct gtctccctgg agttcttgca
                                                                           60
agtcggccag gatgtctcag gctgagtttg agaaagctgc agaggaggtt aggcacctta
                                                                          120
                                                                          180
agaccaagcc atoggatgag gagatgotgt toatotatgg ccactacaaa caagcaactg
                                                                          240
tgggcgacat aaatacagaa cggcccggga tgttggactt cacgggcaag gccaagtggg
atgcctggaa tgagctgaaa gggacttcca aggaagatgc catgaaagct tacatcaaca
                                                                          300
aaagtagaag agctaaagaa aaaatacggg atatganaga ctggatttgg ttactgtgcc atgtgtttat cctaaactga gacaatgcct tgtttttttc taataccgtg gatggtggga
                                                                          360
                                                                          420
                                                                          480
attogggaaa ataaccagtt aaaccagcta ctcaaggctg ctcaccatac ggctctaaca
gattaggggc taaaacgatt actgactttc cttgagtagt tttaatctga aatcaattaa
                                                                          540
aagtggattt tgtaccaaaa aaaaaaaaaa aaaaagtnct gcccggccgg ccntcaaaag
                                                                          600
```

```
gcnaattcan ccccttg
                                                                                 617
       <210> 295
       <211> 606
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (606)
       <223> n = A,T,C or G
       <400> 295
cgaggtactt ttaccatgaa catctctaga ctgtgattat taaatatagt gataatatac
                                                                                  60
atgggtttac tgggatattg aaaaataaaa gataatgaac ccaatttagt aaatcaacat
                                                                                 120
aaatacaaaa cagagcgaat tagcceteta caactgaget cgteetgegt ettgagettg
                                                                                 180
ggttctttct ggaactgtct caaaccttag tgggggaagt gaccttatcc acagattgct
                                                                                 240
tttcccagag gttccgcttg ctggatacgt ctcctggtct caagtcagaa ggtttgggag caggtgactt gtttccatct ggggttttag ttagccattc attgatgccg ctagaaaccc ctaccttcaa gccagcagtt tccttatttg gtgtgcctgc tgcantgggg gatgaaaaca cattcctttc tnccacatac tcttggatgt tgcgtacctg cccnggcngg ccgttcnaaa
                                                                                 300
                                                                                 360
                                                                                 420
                                                                                 480
ggccaattcc acaccactgg cggccgtact aatggatcca aaactcggac cancttggcg
                                                                                 540
natcatnggc atactggttc ctggggnaaa tggattccgt tacattcccc caacttccag
                                                                                 600
ccnggg
                                                                                 606
       <210> 296
       <211> 612
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(612)
       <223> n = A, T, C or G
       <400> 296
ggtacgcggg gtgccagagg aaatcttaaa gcgcctactt aaagaacagc acctctggga
                                                                                  60
tgtagacetg ttggattcaa aagtgatega aattetggae agecaaactg aaatttacea
                                                                                 120
gtatgtccaa aacagtatgg cacctcatcc tgctcgagac tacgttgttt taagaacctg
                                                                                 180
gaggactaat ttacccaaag gagcctgtgc ccttttacta acctctgtgg atcacgatcg
                                                                                 240
cgcacctgtg gtgggtgtga gggttaatgt gctcttgtcc aggtatttga ttgaaccctg
                                                                                 300
tgggccagga aaatccaaac tcacctacat gtgcagagtt gacttaaggg gccacatgcc
                                                                                360
anaatggtee egeaggaagg eegteaagaa nggetegace eggntggtgt tteaaggaag
                                                                                420
aaacattgtg gtcttggtgt ggaaaaaaaa tcantgggcc aactggngga tgaaagacna
                                                                                480
tgccggaana nctgggcttt ggatgacaac ccctgcatgg gcttttgang ccttaccgcc
                                                                                 540
gatccagggt tntnttaaca nggcccggtg gaatgccnaa nccccggtta ctttggagga
                                                                                 600
cccggtnctt qq
                                                                                612
      <210> 297
       <211> 590
       <212> DNA
      <213> Homo sapiens
```

```
<220>
      <221> misc_feature
      <222> (1) ... (590)
      <223> n = A, T, C \text{ or } G
      <400> 297
acgcggggga acacatccaa gcttaagacg gtgaggtcag cttcacattc tcaggaactc
                                                                            60
teettettig ggecaeggaa ttaaceegag caggeatgga ggeetetget eteaceteat cageagtgae cagtgtggee aaagtggtea gggtggeete tggetetgee gtagttttge
                                                                           120
                                                                           180
ccetggccag gattgctaca gttgtgattg gaggagttgt ggctgtgccc atggtgctca
                                                                           240
gtgccatggg cttcactgcg gcgggaatcg cctcgtcctc catagcagcc aagatgatgt
                                                                           300
ccgcggcggc cattgccaat gggggtggaa ttgcctcggg caaccttgtg gctactctgc
                                                                           360
agtcactggg aacaactgga ctcttcngat tgaccaagtt catcctgggc ttcattgggt
                                                                           420
ctgccattgc ggctgcattg cnaggtctac taacttcctg cccttgcctt gcaaaaaaaa
                                                                           480
aaacettgcc agggaaaaag nccccaance ttetgaacca accangggge ccaettttcc
                                                                           540
aaaatacctn gggnggaaaa tncccaattt tgantttcnn aggaaanana
                                                                           590
      <210> 298
      <211> 590
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (590)
      <223> n = A,T,C or G
      <400> 298
ggtactttga gccactctcg catggaaagg agtgtcttta tgcctcgacc tcaagctgtg
                                                                            60
ggetetteca attatgette caccagtgee ggaetgaagt atcetggaag tggggetgae
                                                                           120
ctteeteete cecaaagage agetqqaqae aqtqqtqaqq atteaqaeqa caqtqattat
                                                                           180
gaaaatttga ttgaccctac agagccttct aatagtgaat actcacattc aaaggattct
                                                                           240
cgacccatgg cacatcccga cgaggacccc aggaacactc agacctccca gatttaacta
                                                                           300
aacaaaagaa actetecaec tageactgtt titetteatt gettaetgag agggtttttg
                                                                           360
agaacttaat ctggggggag aactgctttc tcagatcctt aactcccgag aagagaagtc
                                                                           420
cttgtgcaca gaacttgtgg gaaccttcat ccgntgtctt tacctttgga tccagtgtgc
                                                                           480
aagtttcatg acngaatcat taagatatca aatggcctaa tttggngcna atcatggtat
                                                                           540
actgggaaaa ttaggcnaat ggaacttntc accgantttg gtctttaaan
                                                                           590
      <210> 299
      <211> 549
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (549)
      <223> n = A, T, C \text{ or } G
      <400> 299
cgaggtacaa agatctgaca tgtcacccag ggacccattt cacccactgc tctgtttggc
                                                                            60
egecagtett ttgtetetet etteageaat ggtgaggegg ataeeettte eteggggaag
                                                                           120
agaaatccat ggtttgttgc ccttgccaat aacaaaaatg ttggaaagtc gagtggcaaa
```

```
240
gctgttgcca ttggcatcct tcacgtgaac cacgtcaaaa gatccagggt gcctctctct
gttggtgate acaccaatte ttectaggtt ageaceteca gteaccatae acaggttace
                                                                          300
                                                                          360
agtigtogaac ttgatgaaat cagtaatott gocagtotot aaatcaatot gaatggtato
attcacettg atgaggggat cggggtaacg gatggtgcgg gcatcatgag tcaccagatg anggattcct tttgtgccca caaagatctt tctactttgc ancacacact ggcggncgta
                                                                          420
                                                                          480
ctagtggate cacttegnae caacttggeg tateatggge tnactggtne egggggaaat
                                                                          540
                                                                          549
ggtatccnn
      <210> 300
      <211> 591
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(591)
      <223> n = A,T,C or G
      <400> 300
actccagcct gggcgacaga gcaagactcc acctcaaaaa agaaatattt agcaaatatt
                                                                           60
aaaggacaag agggaatatc tgtttaaaaa attataatgc acgttagatg aaaagtaata
                                                                          120
ggatgagatg gttgttgctg aaatagcact tgctatataa attcaaacat tccttttcaa
                                                                          180
attcagette teagaggttt gaetteagat gettgageae tttcaacatt atetttgeet
                                                                          240
ttatccttcn ttatgcggat aaacacaact gctaaaatta taccattgat tttggaaact
                                                                          300
tcccagtcgt tttgtaagct tcactgccga gggaaaatgt aaaatgggga ccccgaaata
                                                                          360
aagtgctgat catcatcaag tagcctcgaa aatgagactt tcaggtgcac tgaaggggat
                                                                          420
ggcagaagaa caagccccgt gtagtccttg ctagcctggg aaggttggca ttcacatcct
                                                                          480
taaggatcan qtqqactttg acnccqaact taaaggaaga acccctatt ntqqqqccac
                                                                          540
cacttgacct tgggccggaa cacccttaag gcgaattcca cacactgggg g
                                                                          591
      <210> 301
      <211> 655
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(655)
      \langle 223 \rangle n = A,T,C or G
      <400> 301
egaggtacte tttaaaaagg gaetgeaggg etgggtgtag tggeteacae etgtaatece
                                                                           60
agcactttgg gaggccaagg caggtgggtc acttgaggcc aggagtttga gaccagcctg
                                                                          120
accaacatgg caaaacccca tctctactaa aatacaaaaa ttagctgggc atgatggtgc
                                                                          180
actcctgtaa tcccagctac ttggtaggct gaagcatgag aattgcttaa acctgggagg
                                                                          240
cagaggttgc agtaagccaa gatcatgcca ctgcactcca gcctgggcaa cagagtaaga
                                                                          300
ctctgtctta ataaataaat aagaaaataa aacggaactg cagtgctaac agtaatttat
                                                                          360
acatttttaa atgttctgag tatgttttga ctgggctagt gtaacaatat actacctga
                                                                          420
aaagtgcagt tttgattgtt ggtggtgtct ttgggtcang aaaagtgaac tgtgccaaga
                                                                          480
agtatttttc aatgacatga atggattnct gttaatgcaa ttgactgaga aaatgngctt
                                                                          540
acgetttett aactgeaaaa agagntttgt ecacateana attgttgaaa etggngetgt
                                                                          600
ttetgttgce tgggatetga tgaetgggat tteetettgg acaaaanace tgatn
                                                                          655
```

```
<210> 302
      <211> 513
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(513)
<223> n = A,T,C or G
      <400> 302
actogtottg gtgagagogt gagotgotga gatttgggag totgogotag gooogottgg
                                                                          60
agttetgage egatggaaga gtteacteat gtttgeacee geggtgatge gtgetttteg
                                                                         120
caagaacaag actotoggot atggagtoco catgttgatg gatootgago ttgaaaaaaa
                                                                         180
actgaaagag aataaaatat ctttaqaqtc qqaatatqaq aaaatcaaaq actccaaqtt
                                                                         240
tgatgactgg aagaatattc gaggacccag gccttgggaa qatcctgacc tcctccaaqq
                                                                         300
aagaaatcca gaaagcctta agactaagac aacttgactc tgctgatttt tttttccttt
                                                                         360
ttttttttta aataaaaata ctattaactg gacttcctaa tatatacttc tatcaagtgg
                                                                         420
aaaggaaatt ccaggcccat ggaaacttgg atatgggtaa attgatgacc aataatcttc
                                                                         480
acttaaagnc atgtcctttg gccgcgaaca cgc
                                                                         513
      <210> 303
      <211> 610
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (610)
<223> n = A,T,C or G
      <400> 303
acgcggggct tgcagagccg gctccggagg agacgcacgc agctgacttt gtcttctccg
                                                                          60
cacgactgtt acagaggtct ccagagcctt ctctctcctg tgcaaaatgg caactcttaa
                                                                         120
ggaaaaactc attgcaccag ttgcggaaga agaggcanca gttccaaaca ataagatcac
                                                                         180
tgtagtgggt gttggacaag tnggtatggn gtgtgctatc agcattctgg gaaagtctct
                                                                         240
ggctgatgaa cttgctcttg tggatgtttt ggaagataag cttaaaggag aaatgatgga
                                                                         300
tetgeagent ggggagetta titetteana cacettnaaa ttgtgggeag atnaagatta
                                                                         360
ttctgtgacc cgtcaattct tanattngta gttggtnact gcatggaatt cngtcagcaa
                                                                         420
gaaangggaa aantcingti caattiggin gnataagaan iggitaatgg tottcaaatt
                                                                         480
cnttattcct tcagancggc caagtacctn ggccgnganc atgcctaagg gctaattcna
                                                                         540
ctcantggng gccgntctan ntggattcca ncttggtacc aancttggng ntattnatgt
                                                                         600
caatanctgg
                                                                         610
      <210> 304
      <211> 596
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(596)
```

<223> n = A, T, C or G

```
<400> 304
ggtacctgga attatctaat tggccagagg tggcgcccga cccatcagtt cgaaatgtag
120
ctaagaagtt ttattctgaa tctgaggaag aggaggactc ttctgatagt agcagtgaca
                                                                          180
gtgagagtga atctggaaag tgaaaagtgg agaacaaggc cgaaagtggg ggaggaagga gacagcaatg aggacagcag tgangactcc tncagtgagc angacagtga gagtggacgg
                                                                          240
                                                                          300
gagtcaggcc tagaaaacan angaacagcc nagangaact caaaagccaa agggaaaaag
                                                                          360
tgattctgaa gatggggaga aggaaaatga aaaatctaaa acttcagatt cttcaaatga
                                                                          420
cgaatctagt tcaattanaa gacagttctt ccgattcttg aatcagaatc agaacctgaa
                                                                          480
agtgaatctt gaatnengaa eagtegetta ggagaaagaa agaaaccaag eaggattgae
                                                                          540
teettttnee aagntgitee tietaaactg gatgatttaa cengniceet caqtqn
                                                                          596
      <210> 305
      <211> 629
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(629)
      <223> n = A,T,C or G
      <400> 305
ggtactttnt tttttttt tttttttt tttttttt tggggattta ntttttattt
                                                                           60
cataatcata aacttaactn tgcaatccan ctaggcatgg gagggaacaa ggaaaacatg
                                                                          120
gaacccaaag ggaactgcag cgagagcnca aanattntng gatactgcga gcaaatgggg
                                                                          180
nggaggggng ctntcctgag ctacaaaagg aatgatctgg tggntaaaat aaaacacaag tcaaacttat tnnagttgtc cacagncagc aatggngatc ttcttgctgg ncttgccatt
                                                                          240
                                                                          300
cctggaccca aagcgctcca tggcctccac aanattcatg ccttctttna ctttgccaaa
                                                                         360
caccacatge ttgccatcca accactcant cttggnagng cagatgaaaa actgggaacc
                                                                         420
atttttnttg ggtccnacat ttccatggca aaangccang accenttget ttaagaagaa
                                                                         480
aatctcatct tcaaattctn ccctaaanga cttgccncan gccntntggg tgngaagcnc
                                                                         540
cccctgncca taaccctgga tatttttgaa agaggancct ntacnaacnt ttttccnggt
                                                                          600
aanaaaaaat tttttntttg acctnccca
                                                                         629
      <210> 306
      <211> 643
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (643)
      <223> n = A, T, C \text{ or } G
      <400> 306
acagggagga atttgaagta gatagaaacc gacctggatt actccggtct gaactcagat
                                                                           60
cacgtaggac tttaatcgtt gaacaaacga acctttaata gcggctgcac catcgggatg
                                                                         120
tectgatece eegegtacat treettgtag actetgttaa treetgeag eteetggttg
                                                                         180
gttctggagc agatgatctc aatgagagag tcctcgtcgg ttcccagccc cttcatggaa
                                                                         240
gcttttatct cagaagcgtc atactgagca ggtgtnttca ataggcccaa aatcaccgtc
                                                                         300
tccaggtggc cagataaggc tgacttcaat gctgatgcaa gntccttttt ggtccttctc
                                                                         360
```

```
tggtaggcga aggnaatate etgtetetgt neattgettg eggntgggca aaatgttgae
                                                                       420
aatqqtqacc tcatccacac ctttgqtctt tgatggntgg ntcaatgttc aaagcatccg
                                                                       480
ctcagcatca aaantaagta tangctttgc agacccatat gcacttgggg gngnngagng
                                                                       540
                                                                       600
acacceteca actgaacttg ccaggatttn tgaaagtaan anttttaaga acttgeegne
                                                                       643
cccanactaa acnnccaatc tagcccnntn cctaacggcc aag
      <210> 307
      <211> 643
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(643)
      <223> n = A,T,C or G
      <400> 307
cgaggtactt ttttttttt tttttttnt ttnttnttnn tttggggatt nantttttat
                                                                        60
ticataatca taaacttaac tetgeaatce aactaggeat gggagggaac aaggaaaaca
                                                                       120
tggaacccaa agggaactgc ancgagagca caaanattct nggatactgc gancaaatgg
                                                                       180
                                                                       240
ggnggagggg tgctctcctn agctacaaaa ggaatgatct ggtggttaan ataaaacaca
agtcaaactt attcnagttn tccacagnca gcaaagggga ncttcttgnt gggcttgcca
                                                                       300
                                                                       360
ticctggacc caaaacgctc catggnetec caaaatttat geettttttt actttgecaa
anaccacatg ctttgccttc caccnctcan tttttgnggg ggnaaataaa aancgggaac
                                                                       420
cnnttgtgtt tggnccnaca ttttccnttg gnaaaaaacc ncgacccctt tntttaagaa
                                                                       480
                                                                       540
naaaatttta nttttaaaat tttcccctaa aaaggactgg cccnaaggcn ttttgggggn
gaagcccncc ntccccnaaa cctggaaaaa ttttggaagc nggacccttt accaaatctt
                                                                       600
                                                                       643
tntcctggtt aaaaaaaat ttttttttt gacctttccc aan
      <210> 308
      <211> 653
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(653)
      \langle 223 \rangle n = A,T,C or G
      <400> 308
                                                                         60
cgaggtacag agagtagett etgtgatgea agaatataet cagteaggtg gtgttegtee
atttggagtt tctttactta tttgtggttg gaatgaggga cgaccatatt tatttcagtc
                                                                        120
agatecatet qqaqettaet ttqeetqqaa aqetacaqea atqqqaaaga actatgtgaa
                                                                        180
                                                                        240
tgggaagact ttccttgaga aaagatataa tgaagatctg gaacttgaag atgccattca
                                                                        300
tacagccatc ttaaccctaa aggaaagctt tgaagggcaa atgacagagg ataacataga
agttggaatc tgcaatgaag ctggatttag gaggettact ccaactgaag ttaaggatta
                                                                        360
cttggctgcc atagcataac aatgaaagtg actgaaaaat ccagaatttc agataatcta
                                                                        420
tctacttaaa catgtttaaa agatggtttg tttgcaagac tttttgcata cttanttcta
                                                                        480
                                                                        540
catgaattaa atcactqqtt tnaaatqaca cttattaatc ctaataactg gtnaacccnc
aaaaaaaaa aaaaaaaa ntacttnccc ggcggccgtc gaanggcaat tcacncctgg
                                                                        600
cggccgtcta tggatccacc cggnccacct gggnaacagg cnactggttc tgg
                                                                        653
```

<210> 309

```
<211> 649
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(649)
<223> n = A,T,C or G
      <400> 309
acttgcaaaa gcacttgaag tcattaaacc agctcatata ctgcaagaga aagaagaaca
                                                                            60
gcatcagttg gctgtcactg cataccttaa aaattcacga aaagagcacc ageggatect
                                                                           120
ggctcgccgc cagacaattg aggagagaaa agagcgcctt gagagtctga atattcagcg
                                                                           180
tgagaaagaa gaattggaac agagggaagc tgaactccan aaagtgcgga aggctgagga
                                                                           240
agagaggctg cgccaggaag caaaggagag agagaaggag cgtatcttac aggaacatga
                                                                           300
acaaatcaaa aagaaaactg toogagagog titggagoag alcaagaaaa cagaactggg
                                                                           360
tgccaaagca ttcaaagata ttgatattga agaccttgag gaaatggatc cagattttat
                                                                           420
catggctnaa cagggtgaac aactggagaa agaaaagaaa gaacttcaga acccttaaga
                                                                           480
atcagaaaag aagattgctn ttttgaagac ccacctttgg aaaaattcct ttgttaagag
                                                                           540
cetttegagg acagaaaatt aagacatggt etggggngee eecqaqqaga aagaattete
                                                                           600
ctgcccttga cgtgaaaggt nttgcataaa atcatgtccn atcttgaga
                                                                           649
      <210> 310
      <211> 319
      <212> DNA
      <213> Homo sapiens
      <400> 310
cgaggtacta gccggacttg gattttctgg aaagatttca gttgaggaac gggaacaaag
                                                                            60
attatgatag ctttccgacc accaccaact tcaatttcct tagctgccgt aatattcagc
                                                                           120
tecetgaget gageettgag gteegagtte atetecaget ecagaagage etgggagatg
                                                                           180
ccggactcga actcgtccgg cttctcgcca ttgggcttca cgatcttggc gctcgaactg
                                                                           240
aacatggett teteetggga gaacttgeeg agegeegget taggaagaga eeeegegtae
                                                                           300
ctgccgggcg ggcgctcga
                                                                           319
      <210> 311
      <211> 646
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(646)
      \langle 223 \rangle n = A,T,C or G
      <400> 311
cgaggtactg atgcaacagt tgggtagcca atctgcagac agacactggc aacattgcgg
                                                                            60
acacceteca ggaagegaga atgeagagtt teetetgtga tateaageae tteggggttg
                                                                           120
tagatgctgc cattgtcgaa cacctgctgg atgaccagcc caaaggagaa gggggagatg
                                                                           180
ttgagcatgt tcagcagcgt ggcttcgctg gctcccactt tgtctccagt cttgatcaag ctgcacatca ctcangattt caatggtgcc cctggagatt ttagtggtga tacctaaagc
                                                                           240
                                                                           300
ctggaaaaaa ggaggtettn tntggeecca aaccaatgtt ctgggetgge caatgaette
                                                                           360
acatggggca atggcaccaa caccggcaga acttgnaccc tattgccaca acatgtcctt
                                                                           420
```

```
atctnaatga nggncttctt tgtgaaaaca aaccccattc cccggaatta agnacaantt
                                                                     480
cttcaaactt gggtggnttc aagggcctcg atngcctgcc catatngggt ttttgccata
                                                                     540
aaacacaacn ttccnnaaag gaatccgant nttgttttgt tggancccat ttttqttccc
                                                                     600
aagaaaattn ggtaatatcc aaattgggga attaggaaaa tgggnt
                                                                     646
      <210> 312
      <211> 622
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (622)
      <223> n = A, T, C \text{ or } G
      <400> 312
cgaggtactt ttgtgagagg gttcaatggg agagctttaa tgcagatgag acttgaagct
                                                                      60
tctgaagaag atctaagtct tgatgaggtt attcaaactc aaatcttgaa tgcataatga
                                                                     120
tgataggcca tggtcttcaa aaacgtggta cttttaatag caacagggtt tcaccatgtt
                                                                     180
ggccaggetg gtctcaaatt cetgacetca agtgatetge ceaettaagt getgggatta
                                                                     240
300
                                                                     360
caatettaae ttaetgnaae eteceettee aggtenaaag aatetttgng etaeeteeta
                                                                     420
natntnggaa tacaagggcg tcccccacct actaattttg ntttttaaga aaaggagggt
                                                                     480
ttancatgtt ggtnngntga teccaacete egacettaan ganeeteege etaattteea
                                                                     540
aaggetggat nttggetgan eccaeceene ttaaccaaaa ttnaaattet tttnteetge
                                                                     600
cgggggcgtt aaagggaatc aa
                                                                     622
      <210> 313
      <211> 674
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(674)
      <223> n = A,T,C or G
      <400> 313
nggacttgaa atcattgaag ttctgcaaaa aggagatgga natgcacaca gaaagaaaga
                                                                      60
tacagaggtc cgcagacggg agctcctaga atccatttct ccagctttgt taanctacct
                                                                     120
gentgaacac geceaagaeg tggtgetaga taagteageg tgtgngtagg tntetgneat
                                                                     180
tccngggaac agacnaattn gaccatnagg naacctgagc ttnccaaagt ncgcaaggct
                                                                     240
gaagaagana ggctneteea ggaageenae gagaaagana aangageegt attttacneg
                                                                     300
aacatgaaca aatcaaaaaa naaaactgtc cgaaaaccgt ttggagcaaa ncaaanaaaa
                                                                     360
cagnacctgg gngcccaaag cattcnaana tatttgttat tancncaccn tgatggattc
                                                                     420
naaacnttat ttttncttgg cncggctggn ccgcccggct ngngnaaaga aaagaacttt
                                                                     480
nctacenete eegaateaag aaaagaanat ggettttttn taaaanneaa eeettgggaa
                                                                     540
aaaattettt gtttaanane eeteeaange eegggaaatt aatteatget ttgtgtgnge
                                                                     600
gacchannaa aaaanaanan ateetteett eeettaann gaaaagggee ttncaaaaaa
                                                                     660
tgattgccca agnc
                                                                     674
```

<210> 314

```
<211> 646
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (646)
      \langle 223 \rangle n = A,T,C or G
      <400> 314
acttttttt tttttttt tttttttt tttgagatgg agtcttgctc tgtcgcccag
                                                                            60
                                                                           120
getggagtge agtgttgega teteagetea ttgeaacete tgeeteecag gtteaagtga
ttctcctgcc tnagcctcct gagtagttgg gactacaggc acatgccacc atgcctggct
                                                                           180
aatttttttg catttttaag tanagacagg gtttcatcat gttggccagg caggtntcaa
                                                                           240
actectgace teaagtgate cacetgtete ageeteecaa agtgetggga ttacaggeat
                                                                           300
gagccactgn accoggocta aaaatgatta cttcttataa aaaggatttc ttccccttca
                                                                           360
caacacttan cttccttttt ctttcctggn aactatgggt ntggngnccg cataaggatc
                                                                           420
tacettnene aagetggaca ntgggggttg etnettgang gnaacteagg ecanataeng
                                                                           480
accetggggg gaacnetaaa ettaettggg tanaaccegg getaacattt etgettgnga
                                                                           540
ngttgattcc ccncaaattt ttaaaaggnn tttcatggcc cttagggcaa ccattttaca
                                                                           600
aagatgggnc acatggnctt ggccgnaacc cctangngaa ttcncn
                                                                           646
      <210> 315
      <211> 666
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(666)
      <223> n = A, T, C \text{ or } G
       <400> 315
                                                                            60
acagtetttg gatatttagg aaggggatgg ggagaaagte agtteteaga acaaattagt
cagetteagt etegteagea gggtetttgg attetttgtt etteegeact tetteaatgt gettateett etetegeaaa egtteeagtt tggeageeat ttgtgeetet eggttetett
                                                                           120
                                                                           180
tattagette catttigtgg gicagttiet ettetgeeat ttiactgaag tigntgttet
                                                                           240
cttctattgc cttctgaagc acttctttct cgtgctctcg tttctcancc agctgcttca
                                                                           300
agacettage tteatgggae ttgegtettt ettetgeage ttetaattte ttetgaattt
                                                                           360
                                                                           420
cctccaggga aagaccttct tctttggaag ggaaaggggg aattctggaa ccagattctt
                                                                           480
ttgacccaag gctgaaaatc agcttaaaag cctggccttg angcacccnt tttcagntct
                                                                           540
ttcacctgga tatcntaaag aagccctngt gattnaaaac aagccnaccg gcantnnatt
ntgncaanan ennggataan gnaateeetg tnaanteena eeeetnaeee eatttteeeg
                                                                           600
ggaccttggc ngnaacccct tanggngaat tcnnccnctn ggcggccgta ctaangggac
                                                                           660
                                                                           666
ccaccq
       <210> 316
       <211> 656
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
```

```
<222> (1)...(656)
      <223> n = A, T, C \text{ or } G
      <400> 316
                                                                           60
actettggtt tgtcaatggg actttccagc aatccaccca agagetettt atccccaaca
                                                                          120
tcactgtgaa taatagtgga tcctatacgt gccaagccca taactcagac actggcctca
ataggaccac agtcacgacg atcacagtct atgcagagcc acccaaaccc ttcatcacca
                                                                          180
gcaacaactc caaccccgtg gaggatgagg atgctgtagc cttaacctgt gaacctgaga
                                                                           240
ttcagaacac aacctactgt gggtgggtaa ataatcagag ccttcccgnc aagtcccagg
                                                                           300
cttgcagctg gccnatgacc aacaggaccc tnactctact tagtgtcaca aggaatgatg
                                                                          360
ganggaccet atgaagtgtg gaaaccagaa ccaattaagt ggtgnccaca cganccagge
                                                                           420
attettgaat ggeeettatg gnecanaaga acceaecatt teeeetnata cacetaatne
                                                                           480
cqtccaqqqt qaaccttaaq ctntctqqca tqcaancctt aaccactqqc aqqattcttq
                                                                           540
qnttaatqaa qqqaacattc nnacccnccc aqaaqttttt attttcaact tacttqqaan
                                                                           600
aacgggggct ntttactgcc ngccataact taacnggggc cnnancggac ttcgnn
                                                                           656
      <210> 317
      <211> 636
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(636)
      <223> n = A, T, C \text{ or } G
      <400> 317
acttttttt tttttttt ttttttttt ttttgnagtca gctatttaat taggttctta
                                                                            60
agacatttag aacaccaatt tgngaggata aattccattc gtcagagcaa acacagatcg
                                                                           120
caggtagece tggagetgag gaatagettt gatttttggt aaaatttgtg agteeacage tttetgatea atettgeget geteegtaat eteatatte eettttetg ggnegaaaan
                                                                           180
                                                                           240
cttacctttc tggggnntgg gcttncgcag cttcttcttn ttgaagtaag catnagtaan
                                                                           300
aagntttggg anttttacan tgntgatann cattttggnt gaagnggnan tgacnaattt
                                                                           360
ctgggggggt cttcgtaaag gaactcnant gaggcccaag ggnccgtccn agtaataagg
                                                                           420
ecetnneane tggttangga aacceeetnt tggeetgggg ggneeangag gntgateeaa
                                                                           480
atggccccgg ggaaaagcng gntcaanttt tnacggctnc tnaaagggtt ttgccnggnt
                                                                           540
taanetttnn ggnentttte agnggaaana cengetttgn nantntaeee eeggnteete
                                                                           600
ggcggaaacc nttagggnna attncncnct gggggg
                                                                           636
      <210> 318
      <211> 654
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(654)
      <223> n = A, T, C \text{ or } G
      <400> 318
cgaggtacgc ggggcctttc tqcccqtqqa cqccqccqaa qaaqcatcqt taaaqtctct
                                                                            60
                                                                           120
cttcaccctg ccgtcatgtc taagtcagag tctcctaaag agcccgaaca gctgaggaag
ctcttcattg gagggttgag ctttgaaaca actgatgaga gcctgaggag ccattttgag
                                                                           180
```

...

```
caatggggaa cgctcacgga ctgtgtggta atgagagatc caaacaccaa gcgctncagg
                                                                            240
ggctttgggt ttgtcacata tgccactgtg gaagaggtgg atgcagctat gaatgcaagg
                                                                            300
ncacacaagg tggatggaag aattgtggaa ccaaagaaaa ctgtcttcag agaagattct taaagaccan gtgcccactt aactgtgaaa aagatatttg gtggtggcat taaagaagac
                                                                            360
                                                                            420
actgaagaac atcactaaga gantattttg aacagtatgg anaaaattgn agngattgaa
                                                                            480
atnatgactg conangoagt ggcancaaan ggggctttgg ctttnnacct ttgacnacca
                                                                            540
tgactenngg ataaaatggn attennaaat ceetentgng aatggeenea etgggaagtt
                                                                            600
ngaaancetn ncaacnagaa agggtnegnt tnnteeneea aangenaang ttte
                                                                            654
      <210> 319
      <211> 659
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (659)
      <223> n = A, T, C or G
      <400> 319
acgcggggaa gccaactcag actcagccaa cagagattgt tgatttgcct cttaagcaag
                                                                             60
agatteattg cageteagea tggeteagae cageteatae tteatgetga teteetgeet
                                                                            120
gatgtttetg teteagagee aaggeeaaga ggeecagaea qaqttqeece aqqeecqqat
                                                                            180
cagctgccca gaaggcacca atgcctatcg ctcctactgc tactacttta atgaaagacc gtgagacctg ggttgatgca anatctctat tgncagaaca tgaattnngg caacctggtg
                                                                            240
                                                                            300
tetgtgetna eccangecca aggtgeeett ggggeeteae tgattaanga aantggeaet
                                                                            360
gatgacttca atggctggaa tggccttcat gacccenaaa aagaacccgc gnttgcactg
                                                                            420
gacagtgggt ccctngntct cttacaagtc tggggcaatt gganccccaa nccatqntaa
                                                                            480
ttenggetae tggggtgage nnaceteage ecaggatttn qaantqqaan geetqnettq
                                                                            540
ggaanacaag ttettettin getngeaagt teaaaaeeta atgeagetgg aaaateatnt
                                                                            600
ctanaactga tcagcattcn accgnttcaa attaaccggc ctttttcant tanttaccg
                                                                            659
      <210> 320
      <211> 664
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (664)
      <223> n = A, T, C \text{ or } G
      <400> 320
ggtactctgc cttttaggag atgaggtaag acatatacat agatggcttt tactagccaa
                                                                             60
ggcaatgtaa atggactaag attotoatgt gacttgaggt tatotgatga atttattoto
                                                                            120
ttcaaaacca cctactttta gagggcatgt ttaacccctc tctttattta aggagggaga
                                                                            180
gaaaaacaca tgtaaccaga attcagagtg ggttactcaa cctaagagaa catacggagt
                                                                            240
tetetttggg aaaacgacaa gactacagtg tteaettege accatgaagt ggeacteetg
                                                                            300
ntatggctgc agantcetet tacttettat gaaaggatge atetgattet gaaattaetg
                                                                            360
atatattega teagttaggg atgntttaaa aagngaaaae caatgeeaca catacaettt
                                                                            420
ctagctttct gaaaatnacc cgacacattn ccnaaaatng agaatttacc ctattacttt
                                                                            480
tagagaaatt tccataatat tcttgggtaa agaancccng ttgggcatat tnccaatttt
                                                                            540
cagnggment ggttttatge cenaganece aataggmtee eccattttt aaggetttt
                                                                            600
```

```
ccacngacga ttttttaaan cnttctnnan tgggggaaga ataatcttaa aagtngnctt
                                                                        660
atnt
                                                                        664
      <210> 321
      <211> 666
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(666)
      <223> n = A, T, C or G
      <400> 321
cgaggtacag tattacagtc agccacagaa gctgtgttgg gggacaagac ccaatccttc
                                                                         60
cccacaccag gcaaagcagt attggacatg agttggcatg tggctgggcc cacqtcctta
                                                                        120
tecceeagge etgaggggag accaeettet gatgataace aacceetage taccaetetg
                                                                        180
tattcatcag gggagggta taaaccccgc atgcaagaag aacccttgcc cccagtgtca
                                                                        240
aatgggatgg ggatgctaga gttatagtaa aggggaaacc ctatgtaagc tgntaacaga
                                                                        300
gttcacaggg gtagggataa cccctgntct tcagctncca aatgngctca ctttccagct
                                                                        360
tetteateeg teateaatge tggeaaagtt teeetnaact gnggeeaggt ttteaegeat
                                                                        420
gggtggctgc acctgggtca aaaaggtggn attggccntt aaggaattag caatcatntq
                                                                        480
ctgggtggga ttccagtgtg taaggaactt anccaactgc atggnttgnt tgtgcanctg
                                                                        540
cttgatggng acaagtttnt gcaccanctn aaggaaggtg gaagcatggg gctcaacctn
                                                                        600
gataagttca tatacttggg geneettget ttgggatetg catntttaca aggnttaten
                                                                        660
tggcan
                                                                        666
      <210> 322
      <211> 631
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(631)
      \langle 223 \rangle n = A,T,C or G
      <400> 322
accggaaagg aagctcccat tcaaaggaaa tttatcttaa gatactgtaa atgatactaa
                                                                         60
ttttttgtcc atttgaaata tataagttgt gctataacaa atcatcctgt caagtgtaac
                                                                        120
cactgtccac gtagttgaac ttctgggatc aagaaagtct atttaaattg attcccatca
                                                                        180
taactggtgg ggtacatcta actcaactgt gaaaagacac atcacacaat caccttgctg
                                                                        240
ctgattacac ggcctggggt ctctgccttc tccccttacc cttccggctc cacccttcct
                                                                        300
gcaacaacag ccctntacct ggggggcttg ntagaagaga tqtqaaqqqt tcaaqqtcqc
                                                                        360
aacctgtggg actactgcta ggtgtgtggg gnggttcgcc tgcacccctg ggttctttaa
                                                                        420
gnettaaagt gatgeeett teeaaceatt attetggnee cacaettete acteeggeet
                                                                        480
tggncnanca taaatgnace cetteaette etntgagaat ggeettegng aagaatenag
                                                                        540
gettteecaa nettetttee eccenttate angggngetg gttttetnet etenaaggte
                                                                        600
ntttgaccgn accaccaaac ttctgaattn t
                                                                        631
      <210> 323
      <211> 647
      <212> DNA
```

```
<213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (647)
      <223> n = A, T, C \text{ or } G
      <400> 323
actgtgggtc gaagtaatgg atacggacgt aaccatcttc gccgccgctg ctgtagctct
                                                                        60
tgccatcagg atggaaggca acactgttga taggtccaaa gtgacccttg actcttccaa
                                                                       120
actettette aaaggeeaaa tggaagaace tggeeteaaa ettgeeaate etggtggagg
                                                                        180
ttgtggttac atccatggct tcctgaccac cgcccaggac cacatggtca tagttggggg
                                                                        240
agagggcage tgagttgaca ggacgttetg teeggaaagt ettetgatgt teaagagttg
                                                                       300
tggagtcgaa aagcttggct gtgttgtcct tggacgcggt cacaaacatg ggcatgtccc
                                                                       360
tggataactg gatgtccgtg atctgcccgg agtgcttctt aacattncca acacctnttc
                                                                       420
aaanttggca ctatactggg tgagctcttc acttttatng gcaacgnatg atcacttccc
                                                                       480
caagggtccc caaacagcac tqqqqaattt agaqncattc caqqqaactt tatqtaqqqt
                                                                       540
tcatggtgca attggttnga tccccaggtc aaaaagttnc aaacactgga nccctttctt
                                                                       600
gtccnnggag aacatgttat ttgccccaag taaaacccng nccggng
                                                                       647
      <210> 324
      <211> 653
      <212> DNA
      <213> Homo sapiens
      <221> misc feature
      <222> (1)...(653)
      <223> n = A, T, C or G
      <400> 324
ggtacttttt ttttttttt ttttttttt ttqaqatqqa qtcttqctct qttqccaqac
                                                                        60
cggagtgcag tggtgcgatc tgggctcact gcaatctcca cctcccgggt tcaagcgatt
                                                                       120
etectgeete ageeteeega gtaaetggga etaeaggtgt gegeeaeeaa geeeagetea
                                                                       180
tttttgtatt tttagtanag atggggtttc acgatgttgg ctaggatggt ctcgatctct
                                                                       240
ggtcagagtc ttttctgtaa aaatccttgg taaagaagca attttagact gtancctgtt
                                                                       300
gcaaatgcnt taaggaagaa gcaaaacaac tgntagtctt tctgaaatga aaaaactacn
                                                                       360
ccagggctgg tatatnnaga gcaaccccaa ccannactnc catcntgatg cccacagggg
                                                                       420
cccactgana naccongaaa angtoonnaa gontaaannt ngangonttg ottttgaaat
                                                                       480
attgcgccng taccnagntn nagacaaacn ngnttaaggc ccnnantntt tggccngant
                                                                       540
ttgcgataaa aaaaacttgg gggtcgctnc nngatcccnn ttgtncccca naanctgggg
                                                                       600
ggatgggttn aagccontgn cnnaaggttt nngtteteec aaggtaaaan nng
                                                                       653
      <210> 325
      <211> 655
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(655)
```

<223> n = A, T, C or G

```
<400> 325
ggtacgcggg gccttttggc tctctgacca gcaccatggc ggttggcaag aacaagcgcc ttacgaaagg cggcaaaaag ggagccaaga agaaagtggt tgatccattt tctaagaaag
                                                                            60
                                                                           120
attggtatga tgtgaaagca cctgctatgt tcaatataaq aaatattgga aagacgctcq
                                                                           180
teaccaggae ceaaggaace aaaattgeat etgatggtet caaggggteg tgtgtttgaa
                                                                           240
agtgagtett getgattige agaatgatga agtigeatti agaaaattea agetgattae
                                                                           300
tgaagatgtt caagggtaaa aactgnctga ctaacttcca tqgcatggat cttacccgtq
                                                                           360
acaaaatgtg gtccatgggc aaaaaatggc agaccatgat tgaagcttac ggtgatgtca
                                                                           420
agactacega atgggtaett gettegtetg gtetggggtg ggtttaetaa aaaaegeaea
                                                                           480
atnanatace gaagaactet tatgettang accaeangte engecaatee ggagaanata
                                                                           540
tggaaatctg accccaaagn gccnaccaat gacttgaaaa annggccatt aaatggttcn
                                                                           600
nacacnttgg aaaagcctta aaaggttgcc aantattaac ctntcatgaa gnttc
                                                                           655
      <210> 326
      <211> 657
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(657)
      <223> n = A,T,C or G
      <400> 326
ggtacgcggg ggaaacggga gtgaacggag agcgtagtga ccatcatgag cctcctcaac
                                                                            60
aagcccaaga gtgagatgac cccagaggag ctgcagaagc gagaggagga ggaatttaac
                                                                           120
accggtccac tctctgtgct cacacagtca gtcaagaaca atacccaagt gctcatcaac
                                                                           180
tgccgcaaca ataagaaact cctgggccgc qtgaaqqcct tcqataqqca ctqcaacatq
                                                                           240
gtgctggaga acgtgaagga gatgtggact gaggtacttt ttttttttt ttnttctttt
                                                                           300
ttttgagata gggneteact gnatnaceca ntntggaatg caattggeat gaacneaget
                                                                           360
tactgnagnc ttccaaacct gggctcaagc aattatnttg nattaacctn ttgagtacct
                                                                           420
gggactntcn cangcaccan ccctqctttq cttacttaaa tttttqtnaa nacnnqqctt
                                                                           480
gettttttte ecaggntggn tenaacteen gaattaaggg atcetteece etcaattttt aaannngetg ngattntnga atangeettt ttgttngeee ttttnacett ttnnnnggtt
                                                                           540
                                                                           600
nnttenngge tttaaneetn eegggggeen tttaaaggng aaatenenee ttggggg
                                                                           657
      <210> 327
      <211> 658
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(658)
      <223> n = A, T, C or G
      <400> 327
ggtacttttt ttttttttt tttttttt ttttttttgg tttgaaacag aaatttattc
                                                                             60
tcanaataat gcacagaagc acaggttgag gctactcttg ggaagcttcc ctccccttcc
                                                                            120
tetteeteet eteeteete tetgaatgee agggagaaca cagttgaagg aaggaaacat
                                                                            180
gcaatcacaa acaatgaaca actntaaaga caaaaaggtt tggtccaaaa gaactcaaca
                                                                            240
taattaatcc aatgactgtg aanagcttca ctgagtagga ttaanatatt gcagatgtan
                                                                            300
ngtttncaca gggtggctnt tcagtgcacc ancggggcct ncttgangga natgaggact
                                                                            360
```

near an area and

```
420
gacncatncg ggaaanatct ttggcctgct tgctaaactt ggggntaaag gcacacnnnc
cgggccaccc gttccactna nngcctgggg accanttgtc aatgnenttt cenaangntt
                                                                         480
titttgntgc cttgtggttg ntittggtti ctggaactgn tcgncctgnc ttgnaaacca
                                                                         540
                                                                         600
ttnttntaac nocttaatgg cotttotttt ennnetggtt ntgnttocaa aatnggatta
ngggttcang ngcccctact tnccgggggc ngttaaangg naattccncc nctggngg
                                                                         658
      <210> 328
      <211> 644
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(644)
      <223> n = A,T,C or G
      <400> 328
acgcggacgg tggtttttgg gcccgtttct gagcagcgct tcctttttgt ccgacatctt
                                                                          60
gacgaggetg eggtgtetge tgetattete egagettege aatgeegeet aaggacgaca
                                                                          120
aqaaqaaqaa qqacqctqqa aaqtcggcca agaaagacaa agacccagtg aacaaatccg
                                                                          180
ggggcaaggc caaaaagaag aagtggtcca aaggcaaagt tcgggacaag ctcaataact
                                                                          240
tagtettgnt tgacaaaget acetatgata aactetgtaa ggaagtteec aactataaac
                                                                          300
ttataacccc agctgtggnc tcttgagaga ctgaagattc naggctncct ggccagggca
                                                                          360
gecetttagg agettettag taaaggaett atnaactggt tttnaancac agaeetcaag
                                                                          420
taattnacac cagaaatncc nnggtggaga atnctccnct gctggtnnag angcatgaat
                                                                          480
aggnncaacc agetntetet gneennacen enettaggne naatteegea eeetgeggee
                                                                          540
gttctnatgg atccnaactn ggtnccaant nggcnnacta tggcatanen tgccctgggg
                                                                          600
aantggttcc nttccaatcc anaanttcta tcgnaactta acgg
                                                                          644
      <210> 329
      <211> 651
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(651)
      <223> n = A, T, C or G
      <400> 329
actattagec atggtcaacc ccaccgtgtt cttcgacatt gccgtcgacg gcgagccctt
                                                                           60
                                                                          120
qqqccqcqtc tcctttqaqc tqtttqcaqa caaggtccca aagacagcag aaaattttcg
                                                                          180
tgctctgagc actggagaga aaggatttgg ttataagggt tcctgctttc acagaattat
tccagggttt atgtgtcagg gtggtgactt cacacgccat aatggcactg gtggcaagtc
                                                                          240
catctatggg gagaaatttg aagatgagaa cttcatccta aagcatacgg gtcctggcat
                                                                          300
cttgtccatg gcaaatgctg gacccaacac aaatggttcc cagtttttca tctgcactgc
                                                                          360
                                                                          420
caagactgag tggttggatg gcaagcatgt ggtgtttggc aaagtgaaaa gaaagggcat
gaatattgtg gaggccatgg aaccgctttg ggtccaggaa tgncnagaac agcaagaaga acaccattgc tgactgngga caactcgaat aagttggact tggggttant ttaaccacca
                                                                          480
                                                                          540
                                                                          600
gaacaattcc tttgtncnta aggagancan ccctcaccca tttgntngca tatcctanaa
actitgggct ttentingti ectitggite aggitteetg giectecane e
                                                                          651
```

<210> 330

```
<211> 643
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (643)
      <223> n = A,T,C or G
      <400> 330
actitintitt tittintnitt tittittitt ciggaaggni cicaggicti tattigcini
                                                                              60
ctcaaattcc aggaatngac ttatttaatt aatccatcaa cctctcatag caaatatttg
                                                                              120
                                                                              180
agaaaacaaa tttatattca gattcttatt ttcagtaggg aagtaagaag ttgcagctca
                                                                              240
ttqcacqtaa aqttqaqaca qanatqqaqa catccagccc cacctntctg gaacaagaaa
                                                                              300
gatgactggg gaggaaacac aggtcancat gggaacaggg gttacagtgg acacaagggn
gggctgnctn ttcacctnct tacattatgc taacagggac ncaaacccat tcaggggcct
                                                                              360
ttgcnaaaag aaatgccaaa agctnttgaa gtcncnaagg ggangcgtga anaaaactgc
                                                                              420
atttnagtee eegggteett ngnegggaae eestnanggn gaaateneea eaetggeggg eegtaetagn ggateeaget nggneecaat tggnggaata tggnnaanae tgtteetgtg
                                                                              480
                                                                              540
ggaaaatggn atccgtccaa ttcnccactt acanneggag cctaaangna aaacntgggg
                                                                              600
ngcctatggg gggctacnnn aataatgggt gcctacggcc cnt
                                                                              643
       <210> 331
       <211> 652
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (652)
       <223> n = A, T, C \text{ or } G
       <400> 331
ggtacagatg gcactgacaa tcccctttct ggtggggatc agtatcagaa catcacagtg
                                                                               60
cacagacatc tgatgctacc agattttgat ttgctggagg acactgaaag caaaatccaa
                                                                              120
ccaggttctc aacaggctga cttcctggat gcactaatcg tgagcatgga tgtgattcaa
                                                                              180
catgaaacaa taggaaagaa gtttgagaag aggcatattg aaatattcac tgacctcagc agcccgattc agcaaaagtc agctggatat tataattcat agcttgaaga aatgtgacat
                                                                              240
                                                                              300
ctccctgcaa ttcttcttgc ctttctcact tggcaaggaa gaaggaagtg gggacagang
                                                                              360
                                                                              420
agatggccct ttcgcttang tggccatggg ccttnctttt cactaaaagg aattaccgaa
cagcanaaag aaagnettga gatagtgaaa atggggatga tatetttaga agggngaaga
                                                                              480
                                                                              540
tggggtggat qaaattattc attcctgnga agnttgnaaa ctgngcgnct tcnnnaaant
                                                                              600
nnnaggcatt centnnetgg centgecatt gecattggnt ceanttgeta tagggatgee
ccttaaancn ntttccnnna anagtnnaaa acttgcnntn ggatccaacc nn
                                                                              652
       <210> 332
       <211> 648
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (648)
```

```
<223> n = A, T, C or G
      <400> 332
cgaggtactt ttttttttt ttttttttt ttttttttgag acagagtttc actcttgtcg
                                                                           60
                                                                          120
cccaggetgg agtgcagtgg cgcgatctcg gtccactgca acctcaccct cccaggatca
agegattete etgeeteage cacetgagta getgggatta caggegeetg ceactacace
                                                                          180
tagccaattt ttgtattttt agaagggaca gcatttcacc atgttggcca ggctggtctc
                                                                          240
gaacteetga teteaggnga teeacecace teageeteee aaagtggngg gattacagge
                                                                          300
gtgagccact gaaagttctc attagttttt tggttaaatt ttaaacataa attatgttat agcaaaaatt cctaagaatt gnaaaccact ttatcagaaa tatcnnaaat tcacaaataa
                                                                          360
                                                                          420
tnccaaaatt tataatagct tttttccaga ctaaaatttt aaagctactg anaagnggna
                                                                          480
aacctnccta nataggattt acctaacatt nnggantaaa aggnanccan ngcctgctaa
                                                                          540
anatccagan tatctaanaa teentneetg nntetennte tatnttttea natecgaatt
                                                                          600
tttgaaccca cnttangata nctnntttcc cccttaacnn taattccc
                                                                          648
      <210> 333
      <211> 656
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (656)
      <223> n = A, T, C or G
      <400> 333
cgaggtacaa gatgtccaaa tattgcgaag atctatttgg ggatctcctg ttgaaacaag
                                                                           60
cacttgaatc acatccactt gaaccaggca gggctttgcc atcccccaat gacctcaaaa
                                                                          120
gaaaaatact cataaaaaac aagcggctga aacctgaagt tgaaaaaaac agctggaagc
                                                                          180
tttgagaage atgatggaag etggagaate tgeeteeea geaaacatet tagaggaega
                                                                          240
taatgaagag gagategaaa gtgetgaeca agaggaggaa geteaeceeg aatteaaatt
                                                                          300
tggaaatgaa ctttctgctg atgacttggg tcacaaggaa gctgttgcaa atagcgtcaa
                                                                          360
gaaggettea gatgaeettg aacatgaaaa caacaaaaag ggeetggtea etgtagaaga
                                                                          420
tgagcangcg tggatggcat cttataaata tgtaggtgct ccactaatat ccatncatat
                                                                          480
ttgtcaccat gatcaactac cccagnctgt naaggttcaa ggtttcatgt ggcanaagaa
                                                                          540
cccatattc ttttacatgg cttctttaat gaatcatcgg cttggtactg aaaccctgcc
                                                                          600
attgaattgc attntacaac ggcaatgagc natttcccca gggaggccng cnttct
                                                                          656
      <210> 334
      <211> 647
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (647)
      <223> n = A, T, C or G
      <400> 334
acgcgggcgg gaagtgcaga ggcaaatqca tttagtgttc ttcagcatgt cctcggtgct
                                                                           60
gggccacatg tcaagagggg cagcaacacc gccagccatc tgcactaggc tgttgccaag
                                                                          120
gcaactcagc agccatttga tgtttetqca tttaatgcca gttactcaga ttctggactc
                                                                          180
tttgggattt atactatete eeaqqeeaca qetqetqqaq atqttateaa qqetqeetat
                                                                          240
```

```
aatcaagtaa aaacaatage teaaggaaae ettteeaaea eagatgteaa getgeeaaga
                                                                              300
acaagetgaa agetggatae etaatgteaa tggagtette tgagtgntte etggaagaaa
                                                                              360
gtcgggtccc aagctctaag tgctggntct tacatgccac cattcacaag tctttaacag aatgattcan tggctaatgc tgatatcata aatgcgnaaa naaagtttgg ttctggcnag
                                                                              420
                                                                              480
aagtcaatgg cancaagtgg naaatttggg acatacnent ttgtgataag tggaataetg
                                                                              540
gngcncnctt acngganana cttaacgttn tttaanccaa acacaaccct tqaaaqnnna
                                                                              600
agctctaaan accattggct tttttcnggg ngnaaaaaag gcttaag
                                                                              647
       <210> 335
       <211> 657
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (657)
       <223> n = A,T,C or G
       <400> 335
acaggicaga gicticitti citticitti tgagatggag tcitgcictg tigccagact
                                                                               60
ggagtgcagt ggtgcgatct gggctcactg caatctccac ctcccgggtt caagcgattc
                                                                              120
tectgeetca geeteeegag taactgggae tacaggtgeg egecaccaag cecagetcat
                                                                              180
ttttgtattt ttagtagaga tggggtttca cgatgttggc taggatggtc tcgatctctg
                                                                              240
gtcagagtct tttctgtaaa tatccttggt aaagaagcaa ttttagactg tagctgttgc
                                                                              300
aaatgettta aggaagaage aaaacaactg teaagtette etgaaatgaa gaaactacae
                                                                              360
cagggetget atateagage aaccecaace ageaetteaa teatgatgee nacagtggee
                                                                              420
cagctgagag accnggagaa agttccagat gcanagactg ngatgctctt gactatggaa
                                                                              480
atattgcggc cagtaccaag ttagagacca aacaggcata ngnncccgta ttaattgqcq
                                                                              540
tgaattttgc gataaganaa cttqqqqqqq tqctqcqqat nccatqatcn ccaqaaaact
                                                                              600
tnngggatgg ggtanaggcc catggcagaa aggttanggt ccttccaaag naaaana
                                                                              657
       <210> 336
       <211> 649
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (649)
       <223> n = A, T, C or G
       <400> 336
ggtacgcggg caactatgga attccacage gtgctctgcg gggtcactcc cactttgtta gtgatgtggt tatctcctca gatggccagt ttgccctctc aggctcctgg gatggaaccc
                                                                               60
                                                                              120
tgcgcctctg ggatctcaca acgggcacca ccacgaggcg atttgtgggc cataccaaqq
                                                                              180
atgtgetgag tgtggeette teetetgaca aceggeagat tgtetetgga tetegagata
                                                                              240
aaaccatcaa gctatggaat accctgggtg tgtgcaaata cactgtcagg atgaaaacca
                                                                              300
cttaaantgg gtgnettgng necettning eccaacagea acaaccetat tategiette
                                                                              360
tgnggctggg acaaactggn taaaggatgg aacctggcta actgnaagct gaaaaacaac
                                                                              420
cacattgggc acacangcta tntgaacacc gngactggct ttttcagang gatcctntgn
                                                                              480
gettntggag geaaggatgg geaageeatg ttatnggaae tenaceaang caacacettt cacetttaan ggggggacat tatnaaegee ttgggttaae ettaaegttn ttggtttgng
                                                                              540
                                                                              600
ctgcncaggc ccacattaaa aatgggattt aanggaaana catttnann
                                                                              649
```

```
<210> 337
       <211> 652
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (652)
       <223> n = A, T, C \text{ or } G
       <400> 337
actettggtt tgtcaatggg actttccage aatecaceca agagetettt atececaaca
                                                                                60
tcactgtgaa taatagtgga tcctatacgt gccaagccca taactcagac actggcctca ataggaccac agtcacgacg atcacagtct atgcagagcc acccaaaccc ttcatcacca
                                                                              120
                                                                              180
gcaacaactc caaccccgtg gaggatgagg atgctgtagc cttaacctgt gaacctgaga
                                                                              240
ttcagaacac aacctacctg tggtgggtaa ataatcaaga gccttccggt cagtcccagg
                                                                              300
ctgcagctgt caatgacaca ggaccctnac tctactcagt gtcacaagga atgatgnaag
                                                                              360
gaccctatga atgtggaatc cagaacgaat taaagcgttg accacagcga ccangcatcc
                                                                              420
tgaatgeett tttgggeean acgaccccac catttteccc teataccact attaccgtee
                                                                              480
aggggtgnac cttagncttt tcttgccatg cagcctttac ccaccttgac agnattcctg
                                                                              540
gctggatgtt gggaacatna gnacncacnc aagagetntt ttttccaaca tnatgggaaa
                                                                              600
acanngnnet tatactgeag gecattaett ngeentngee eagnnggetn eg
                                                                              652
       <210> 338
       <211> 651
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (651)
       <223> n = A, T, C \text{ or } G
       <400> 338
ggtacatttg aacacacggc tgtgttaaag atgctgctaa tgtcagtcac tgggtgcact
                                                                               60
aaaggatete ttattttatg taaaacgttg ggattgacaa gatagatetg acaetetgtt
                                                                              120
aagttaccct ctgaagctac ttcttgtgaa atactaatga cagcatcatc ctgccaagcg
                                                                              180
aaagaggcag gcataagcaa ggacaaatta aaagggggta agagccttat catgatgagg
                                                                              240
agtettgntt tgacatettg ggaaaagetg ceatagtgtg aaagtegtea attteteace
                                                                              300
atggtttgca gtttgactgn ctctagttag ggtgaagtct ctgagtggca cacaccttaa gcctgaaggn tttcccttta aattttcatt gagttggccc tcttcagcat atanggcttt
                                                                              360
                                                                              420
aagaacagaa canaccttgg ttttaagtgg gtccatggga taaaatggga atggangact
                                                                              480
ngaagaattc aagggctggg ccatctngca gtattctgaa tatcgaaaat ncnccaaggc
                                                                              540
tgctatataa anccccctgg gcaanacttc aatcggaanc ccacggnggc ccnactnana
                                                                              600
gncaggaccn ttccaantgg aacatnggan tggggccttt gaggcnnggn n
                                                                              651
      <210> 339
      <211> 634
      <212> DNA
      <213> Homo sapiens
      <220>
```

```
<221> misc feature
       <222> (1) ... (634)
       <223> n = A, T, C or G
       <400> 339
actttttttt tttttttt ttttttctag tttcagttat ttattgattt aatcattgta
                                                                               60
 atctccaata gagattacaa tagagatctc caacatgatt tcatgcattt agaggagaaa
                                                                              120
tatttcctgg ttaagtggaa aattgtgcgg atgtggcttc tggaanacct tcattctaaa
                                                                              180
gcagcgttat agtgaaacat ttcatttana aatctggacg ttccttcttc agcttgctgt
                                                                              240
aatccacatt cactgagtag aacttgtatt gatcattggg acccagtttg ttccagggct
                                                                              300
ctgggttatt tctgcccaac aaacatctgg attgaacaat gccagacgca agagatcagt gttgctccag tagctccagt tccaataaat acnaagaggg ggatcaaagc tcggatgctt
                                                                              360
                                                                              420
cttggcctga ccgatgatct ggcggancat gtttgcngca aagtctccga ctggaaagga
                                                                              480
ganaaccgcc taccccaagc cctaagctaa aaattatntg ccccgcgacc ttggncgnga
                                                                              540
ccenctaagg caattecace actggeggee gtetaangga tecaettggg ccaaettgng
                                                                              600
naacatggca nactggtcct ggggaangta tccc
                                                                              634
       <210> 340
       <211> 655
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (655)
       \langle 223 \rangle n = A,T,C or G
       <400> 340
ggtactcttc cactcaggta tccgtgcggc cactccagca cacgcagtat gagcgcttca
                                                                              60
tecectegge etacecetae tacgecageg cettetecat gatgetgggg etetteatet
                                                                             120
tcagcatcgt cttcttgcac atgaaggaga aggagaagtc cgactgaggg gctagagcc
                                                                             180
tctccgcaca gcgtggagac ggggcaggga ggggggttat taggattggt ggttttgttt
                                                                             240
tgctttgttt aaagccgtgg gaaaatggca caactttacc tctgtgggag atgcaacact gagagccaag gggtgggagt tgagataatt tttatataaa agaagttttt ccactttgaa
                                                                             300
ttgctaaaag tggnattttt cctatgtgca gtcactcctc tcatttctaa aatagggacg
                                                                             360
                                                                             420
tggccaggca ccgtggctca tgcctgtaat ccacactttt ggaggncnng caagcggtta
                                                                             480
cgaagtcagg agatcgagac tattctggtt acacgnaaaa cctgncttac taaaagtacc
                                                                             540
tgcccggccg gccgntcaaa ggcgaatcca cacactggcg ggcgtactan tggatnccaa
                                                                             600
cttggaccaa cttggngnaa tatggcatac tggttcctgg nggaaatggt accnn
                                                                             655
       <210> 341
       <211> 648
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (648)
       <223> n = A, T, C or G
       <400> 341
acgaacctac agttttaact gtggatattg ttacgtagcc taaggctcct gttttgcaca
                                                                              60
gccaaattta aaactgttgg aatggatttt tctttaactg ccgtaattta actttctggg
                                                                             120
```

```
ttgcctttgt ttttggcgtg gctgacttac atcatgtgtt ggggaagggc ctgcccagtt
                                                                      180
gcactcaggt gacatcctcc agatagtgta gctgaggagg cacctacact cacctgcact
                                                                    . 240
aacagagtgg ccgtcctaac ctcgggcctg ctgcgcagac gtccatcacg ttagctgtcc
                                                                      300
cacatcacaa gactatgcca ttggggtaag ttgtgtttca acggaaagtg ctgtcttaaa
                                                                      360
ctaaatgtgc aatagaaggn gatggtgcca tectacegne tttteetggt tectanetgn
                                                                      420
gtgaatacct gctacgtcaa atgcntacca ggttcattct ncctttnact aaaacacaca
                                                                      480
ggtgcaacag acttgaatgc taagtatacc taattggata tgggatttaa ttttctttct
                                                                     540
tacaancatt tgtattgcta acaggccaaa atttcagtta cccttagggt ggttaacaat
                                                                     600
cnaattaaac ctgggaggca tacnttgnct aaatattact gnaaaaaa
                                                                     648
      <210> 342
      <211> 342
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (342)
      <223> n = A,T,C or G
      <400> 342
60
tggctntana gggggtanag ggggtgctat agggtaaata cgggccctat ttcaaanatt
                                                                     120
tttaggggaa ttaattctag gacnatgggc atgaaactgn ggtttgctcc acanatttca
                                                                     180
nagcattgac cgtagtatac ccccggtcgt gtancggtga aagtggtttg gtttaaacgt
                                                                     240
ccgggaattg catcigittt taagcctaat gtggggacag ctnatgagtg caaaacgtct
                                                                     300
tgngatgtaa ttattatacc aatgggggct ttaatcggga at
                                                                     342
      <210> 343
      <211> 484
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (484)
      <223> n = A,T,C or G
      <400> 343
ggtacgatgc ctagtgatga gtttgctaat acaatgccag tcaggccacc tacggtgaaa
                                                                      60
agaaagatga atcctagggc tcagagcact gcagcagatc atttcatatt gcttccgtgg
                                                                     120
agtgtggcga gtcagctaaa tactttgacg ccggtgggga tagcgatgat tatggtagcg
                                                                     180
gaggtgaaat atgetegtgt gtetaegtet attectaetg taaatatatg gtgtgeteae
                                                                     240
acgataaacc ctaggaagcc aattgatatc atagctcaga ccatacctat gtatccaaat
                                                                     300
ggttcttttt ttccggagta gtaagttaca atatgggaga ttattccgaa cctggtagga
                                                                     360
taagaatata aacttcaggg tgaccgaaaa atcagaatan gtgttggtat agaatggggt
                                                                     420
ettettette ngeggggten aanaaggtgg tggtneegeg teetggeeng genggegete
                                                                     480
gaan
                                                                     484
     <210> 344
     <211> 657
     <212> DNA
     <213> Homo sapiens
```

```
<220>
       <221> misc feature
       <222> (1)...(657)
      \langle 223 \rangle n = A,T,C or G
cgaggtacgc gggattgttc tggggcttgt cctcctttct gttacggtcc agggcaaggt
                                                                         60
ctttgaaagg tgtgagttgg ccagaactct gaaaagattg ggaatggatg gctacagggg
                                                                        120
aatcagccta gcaaactgga tgtgtttggc caaatgggag agtggttaca acacacgagc
                                                                        180
tacaaactac aatgetggag acagaagcac tgattatggg atatttcaga tcaatagceg
                                                                        240
ctactggtgt aatgatggca aaaccccagg agcagttaat gcctgtcatt tatcctgcag
tgctttgctg caagataaca tcgctgatgc tgtagcttgt gcaaaaangg ttgtcccgtg
                                                                        300
                                                                        360
atccacaagg cattaagagc atgggtggca tggagaaatc gttgtcaaaa cagagatgtc
                                                                        420
cgcagtatgt tcaanggtgt ggagtgtaac tncagaattt tccntcttca ctcatttggc
tetetacatt aaggagtagg aaataantga aaggteeet ceattaattt eeetteaaca
                                                                        480
                                                                        540
aataattttt tccgaaacng gaccaaatat ggccttcttn tagannataa tgtcntaagg
                                                                        600
ggnatttatt ttaagcnnca canttttaat ttgcaaatna ctatctgggg aaaatac
                                                                        657
      <210> 345
      <211> 662
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (662)
      <223> n = A, T, C or G
      <400> 345
ggtacgcggg cgactcttag cggtggatca ctcggctcgt gcgtcgatga agaacgcagc
tagetgegag aattaatgtg aattgeagga cacattgate ategacaett egaaegeaet
                                                                         60
                                                                        120
tgengeceeg ggtteeteee ggggetaege etgtetgage gtetettgea aaaaaaaat
aaannanaan acancaagta caatttaatg cntanaaagg cctctctcca taaaactcan
                                                                        180
cnctttacag atgtangaat atataagcnn tgccaaaatt actaatntgc cacatacnna
                                                                        240
gcatcaattc caggtgctag tnagngggaa aaaaanttgg agaattcggc cctcgangag
                                                                        300
                                                                        360
ctccanannt taanctnoct tactaantno canggttott toaagcatgg aaaaattaat
ngtgctncat ngatnaangn cttgtcattg ggccttnttt cctngacctg gcccggccgn
                                                                        420
                                                                        480
ccgttcnaaa ggctaaatcc agacactgcg gccgttntaa tggttcnnac ttgggccaag
                                                                        540
cttgggnaat catgggcaaa gctgttcctg ggnnaaatnt tatccnctcc aattcncaca
natacgaanc tgaancttaa gtgtnanntn gggngctaaa agtggcnaan ctcccttnat
                                                                        600
                                                                        660
99
                                                                        662
      <210> 346
      <211> 654
      <212> DNA
      <213> Homo sapiens
      <221> misc_feature
      <222> (1) ... (654)
      <223> n = A, T, C or G
```

```
<400> 346
acttettgge egecteacta geacteteeg eetgettttt aaaggettea ttggaggeea
                                                                           60
gcagcgtggc ctgctgcgaa atgagagtca ccaggcgtct aagcaggaag gacagcagcg
                                                                          120
aggaaaagcc agcaatgtag agattcctct gggcacggaa aagcttcatg tggaagtgct
                                                                          180
ccatggcccc gggattgttc tggaggttca ccttttccgt cacatcatca tacttccgaa
                                                                          240
tttcgcgcac ggcatcgatg accaacagca caaggatgac aatgagaacc acaaagaagg
                                                                          300
tgttgccata ggacactaac aactccacca gccgggactt gaaaatcttc tgccatcttt
                                                                          360
taggagaaat gaagggaatg cagagaagca acacaacaaa gaccttcgca tagaggaagg
                                                                          420
tggcaactgc agtccactgc agactcatcc tggtgctana agggttccac aggaagatgt
                                                                          480
gaacttgttn cgagtttcca cagtcaacgt gtcccccgta ccttnggccg ngaacacnct
                                                                          540
taaggcgaat tecaccaetg enggeegtet antggateea aetnggneea aettggegaa
                                                                          600
tatggcaaat tgttctnggg naaatggttc ngtcaattcc ccantacnac cgga
                                                                          654
       <210> 347
       <211> 536
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (536)
      <223> n = A, T, C or G
      <400> 347
ggtactaatt taaggtaaca attctcgagg taaaataagg cattatagta acacaatttt
                                                                           60
catgcctcag caattaacaa tgattttcgt ttaattctct tccaactcta cagacataat
                                                                          120
tctgctttca ccttcatcac gctttcatat ggttttaaca ggggatacac ctcctcttct aagaatctct gcacctgctg ggaggcacga ccagtgaaag aagaaggatc cagtaaatga
                                                                          180
                                                                          240
tccaactggg agtgaatggg actgaagtag gcatcaacct ggatacgctc tatgaggnca
                                                                          300
ttgcaccccc ttcctgctta accacagaag ctgcctgctg agaaagcact ctgattttct
                                                                          360
catggcaatc ctggcggcta ccttcacttt gaccatggcc atgatgatgg tctctgtggc
                                                                          420
catgaaange agetettgee gaatgegeeg teaattaett tggggtaeet geeenggeeg
                                                                          480
geegntenaa nggegaattt cagecaetgg engnegtaet agnggateea aetegg
                                                                          536
      <210> 348
      <211> 665
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(665)
      <223> n = A,T,C or G
      <400> 348
ggtacgcggg gagtcggcgt aggcettagg tgggttcgtg cgccttctac ctcgctgttt
                                                                          60
eggtttteet ggeteetegg ceettttete eeetgttgea getgggageg gaegaagege
                                                                          120
gaagetggga tittttacig teteetgaag aatttaacae aaacatggat atcagaccaa
                                                                          180
atcatacaat ttatatcaac aatatgaatg acaaaattaa aaaggaagaa ttgaagagat
                                                                          240
ccctatatgc cctgttttct cagtttggtc atgtggtgga cattgtggct ttaaagacca
                                                                          300
tgaagatgag ggggcaggcc tttgtcatat ttaaggaact gggctcatcc acaaatgcct
                                                                         360
tgagacaget accaggattt ccattttatg gtaaaccaat gccaatacag tatgcaaaaa
                                                                         420
cagattcgga tataatatca aaaatgcgtg gaacttttgc ttaaaaaaaa aaannnnnna
                                                                         480
```

 $t_{r-r}$ 

```
naaaaaagte etgeenggee geeegtteaa anggegaatt naccaetgge ggeegtteta
                                                                        540
gnggatccaa ctnggnacca acttggcgta atatggcaaa actggtnccg ngngaaatgg
                                                                        600
tatecgttan aatteccaca etteaacegg aacetnaang taaacetggg geetaagagn
                                                                        660
gacnn
                                                                        665
      <210> 349
       <211> 474
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(474)
      <223> n = A, T, C or G
      <400> 349
acttcgtcag tttgtaagac atgagtccga aacaactacc agtttggttc ttgaaagatc
                                                                         60
cctgaatcgt gtgcacttac ttgggcgagt gggtcaggac cctgtcttga gacaggtgga
                                                                        120
aggaaaaaat ccagtcacaa tattttctct agcaactaat gagatgtggc gatcagggga
                                                                        180
tagtgaagtt taccaactgg gtgatgtcag tcaaaagaca acatggcaca gaatatcagt
                                                                        240
attccggcca ggcctnagag acgtggcata tcaatatgtg aaaaaggggt ctcgaattta
                                                                        300
tttggaaggg aaaatagact atggtgaata catggataaa aataatgtga ggcgacaagc
                                                                        360
ncaaccatca tagettgatn atattatatt tetgagtgee agaccaaaga gaaggagtnt
                                                                        420
aaanggatga tentettttg ggeateattt tgggacettn ggeegggaac acce
                                                                        474
      <210> 350
      <211> 452
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(452)
      <223> n = A, T, C or G
      <400> 350
acgcggggac cgtggagagc agagcgcggc ggctggaagc tgctaagtca gagccgcgat
                                                                        60
gtteeggatt gagggeeteg egeegaaget ggaeceggag gagatgaaac ggaagatgeg
                                                                       120
cgaggatgtg atctcctcca tacggaactt tctcatctac gtggccctcc tgcgagtcac
                                                                       180
tccatttatc ttaaagaaat tggacagcat atgaagacag gacatcacat atgaatgcac
gatatgaaga geetggttae agtttegaet eetetetgea agtgaatagg eecagaaagg
                                                                       240
                                                                       300
tgtaagagac tetttgaatg gacataaaat tetgettgtt aagaacaagt ttggetetgg
                                                                       360
taactgacct tcaaagctaa aatataaaac tatttgggaa agtatgaaac gatgtcttcg
                                                                       420
tgatctggtg taccttggnc gngaccacgc tt
                                                                       452
      <210> 351
      <211> 616
      <212> DNA
      <213> Homo sapiens
      <221> misc_feature
      <222> (1) ... (616)
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<223> n = A, T, C or G<400> 351 ggtacgcggg aataattcca tagtcaagag catcacagtc tctgcatctg gaacttctcc 60 tggtctctca gctggggcca ctgtcggcat catgattgga gtgctggttg gggttgctct 120 gatatagcag ccctggtgta gtttcttcat ttcaggaaga ctgacagttg ttttgcttct 180 teettaaage atttgeaaca getacagtet aaaattgett etttaceaag gatatttaca 240 gaaaagacte tgaccagaga tegagaceat ectagecaae ategtgaaae eccateteta 300 ctaaaaatac aaaaatgage tgggettggt ggegegeace tgtagteeca gttactnggg 360 aggetgagge aggagaatng ettgaaceeg gnaggtggag attgeagtga geeagategn 420 achaetgnae teagtetgge aantgagnag getteeatet nanaangaan aganangang aetntnaeet ggaeetgeen ggeeggtegt ttgngeaggt enggagattt atteeettng 480 540 ggtggggngc nntaattggn tgntgggccn attcangttt tgggaatttc nncttggnnn 600 naaaanggga aatttt 616 <210> 352 <211> 603 <212> DNA <213> Homo sapiens <220> <221> misc\_feature <222> (1) ... (603) <223> n = A,T,C or G<400> 352 ggtacggcac ttggcgtaaa gccgcttccc tcaagagtaa ctacaatctt cccatgcaca 60 agatgattaa tacagatett ageagaatet tgaaaageee agagateeaa agageeette 120 gagcaccacg caagaagatc catcgcagag tcctaaagaa gaacccactg aaaaacttga 180 gaatcatgtt gaagctaaac ccatatgcaa agaccatgcg ccggaacacc attettcgcc 240 aggccaggaa teacaagete egggtggata aggeagetge tgcageageg geactacaag 300 ccaaatcaga tgagaaggcg gcggttgcag gcaagaagcc tgtggtaggt aagaaaggaa 360 agaaggetge tgttggtgtt aagaageaga agaageetet ggtgggaaaa aaggeageag 420 ctaccaagaa aaccagccc tgaaaagaac ctgcagagaa gaaacctact acngaggaga agaagcctgc tgcataactc ttaaatttga atatttcntt aagggcnaat nttttggcag 480 540 gttetttgga taagaentnt tttengngtg ggaaaataan tnnnntattn nnggetntee 600 tgg 603 <210> 353 <211> 604 <212> DNA <213> Homo sapiens <220> <221> misc\_feature <222> (1) ... (604) <223> n = A, T, C or G<400> 353 ggtaccgact gtttttgaca actatgcagt cacagttatg attggtggag aaccatatac 60 tettggaett tttgataetg cagggeaaga ggattatgae agattaegae egetgagtta 120 tccacaaaca gatgtatttc tagtctgttt ttcagtggtc tctccatctt catttgaaaa 180 cgtgaaagaa aagtgggtgc ctgagataac tcaccactgt ccaaagactc ctttcttgct

240

```
tgttgggact caaattgatc tcagagatga cccctctact attgagaaac ttgccaagaa
                                                                        300
 caaacagaag cetateacte cagagactge tgaaaagetg geeegtgace tgaaggetgt
                                                                        360
 caagtatgtg gagtgttctg cacttacaca gaaaggccta aagaatgtat ttgacgaagc
                                                                        420
 aatattggct geeetggace tneagacega agaagaeece aagtgtgtge tgetatgaae
                                                                        480
 atctttcaga gcctttcttg nacagctgga ttggcatctt cttaaagcca tgnttaaatt
                                                                        540
 caacttanga ttaaaattaa aattcgtttt gcannatggc caatgcctgg actaacccan
                                                                        600
 ggcn
                                                                        604
       <210> 354
       <211> 631
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(631)
       <223> n = A, T, C or G
       <400> 354
ggtacttttt ttttttttt ttttttttt tttgggacgg agtcatgctc tgtcgcccag
                                                                         60
gctggagtgc agtggcatga teteggetca etgcaagete egceteeegg geteatgeca
                                                                        120
tteteetgee teageeteee gagtagetga gattatagge acetaceaee acgeeegget
aatttttgta tttttagtag agacggggtt tcaccatgtt gaccaggctg gtctcgaact
                                                                        180
                                                                        240
cetgacetta ggtgatecae tegeetteat etcecaaagt getgggatta caggegtgag
                                                                        300
ccaccgtgcc tggccacgcc caactaattt ttgnattttt agtaagagac agggtttcac
                                                                        360
catgttggcc aaggctgctc tttgaactcc tgacctcatg taatcgacct gcctttggcc
                                                                        420
ttccaaaagt getgggatta ccaggtgtga geccacaage ceeggnacet ggeenggeng
                                                                        480
gccgtttaaa agggcgaatt cagcacaatg gnnggccgta ctaaggggat ncnanctttg
                                                                        540
nanccaactt tgggggaaat atggggcana actggttcct ngngnaaatg gtaaccgtta
                                                                        600
caaattcccn caaanttttg nnccgggagg n
                                                                        631
      <210> 355
      <211> 626
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (626)
      <223> n = A, T, C \text{ or } G
      <400> 355
ggtacgatgc ctagtgatga gtttgctaat acaatgccag tcaggccacc tacggtgaaa
                                                                         60
agaaagatga atcctagggc tcagagcact gcagcagatc atttcatatt gcttccgtgg
                                                                        120
agtgtggcga gtcagctaaa tactttgacg ccggtgggga tagcgatgat tatggtagcg
                                                                        180
gaggtgaaat atgetegtgt gtetaegtet attectaetg taaatatatg gtgtgeteae
                                                                        240
acgataaacc ctaggaagcc aattgatatc atagctcaga ccatacctat gtatccaaat
                                                                        300
ggttcttttt ttccggagta gtaagttaca atatgggaga ttattccgaa gcctggtagg
                                                                        360
ataagaatat aaacttcagg gtgaccngaa aaatcagaat aggtgtttgg tttagaatgg
                                                                        420
ngtettetne ttengetggg gtnnaagaan gtnggggtte nngegtnetn gntegggegg
                                                                        480
ntggttttaa nggccnaaat tcnngnataa ttggeggeng ttactaagng gnatctanct
tggtnccaaa nttggngnta atcatggtnc tagctngtnc tcngtgntaa attggntncc
                                                                        540
                                                                        600
tgttaaattn tntnnaatnt tntggc
                                                                       626
```

```
<210> 356
       <211> 617
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (617)
       <223> n = A,T,C or G
       <400> 356
actitititt tittititt tittiticta gittcagita titatigati taatcatigi
                                                                         60
aatctccaat agagattaca atagagatct ccaacatgat ttcatgcatt tagaggagaa
                                                                        120
atatttcctg gttaagtgga aaattgtgcg gatgtggctt ctggaanacc ttcattctaa
                                                                        180
agcagcgtta tagtgaaaca tttcatttan aaatctggac gttccttctt cagcttgctg
                                                                        240
taatccacat tcactgagta naacttgtat tgatcattgg gacccagttt gttccagggc
                                                                        300
tetgggttat ttetgtecca acaaacatet ggattgaaca atgceagacg caagagatae
                                                                        360
agtgttgctc cagtagctcc agttccaata aatacnaaga gggggatcaa gctcggatgc
                                                                        420
ttcttggcct gaccgatgat ctggccggaa ncatgtttgc cggcaaaagg ctccnacttg
                                                                       480
ggaaagggga naacccgcct aaccnccagg gcctaagctt aaaatttttg gccccgggta
                                                                       540
cettggeegg gaccceetaa gggngnaatt cenneceett ggggggeegt ttaangggan
                                                                       600
ccaacttggn ccaaatt
                                                                       617
      <210> 357
      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (611)
      <223> n = A,T,C or G
      <400> 357
ggtacttttt ttttttttt tttttttt ttttaggcaa agaactttat taatctttgt
                                                                        60
ttcaaacttg attcccaggc ttcttcggct taattagctg caaagaatga attgngtata
                                                                       120
agcaaaaact gaaaagagct gcagtgtcca aggggcttgg gcttaaaaat attagagatc
                                                                       180
tagattttat cagatccata aacaaaaatt tottaaaaag cagtcataat ataaaatagc
                                                                       240
ageteceagt aaettettea ggntttatet teagaagttg aeteaattea gtttgeetea
                                                                       300
ttcttggaag cctcatcaaa attctccaca agatctggaa cttcatcatc atcatcctct
                                                                       360
ccagtaacaa gtggngcttt tccatcccca gantggttgg gcanaacttt ngnccagctc
                                                                       420
cttaacttag cagactattc ggacccaagc inggiinaaa aanctgggaa cnattintgn
                                                                       480
naactggttt ggttnaacan ggcntgnaag ggggaaaggn gtnccctgcc caaaaaaccn
                                                                       540
ggacctttag ggtgnnaaag gggacctggc cctgggttgg aaccaanten cettttnana
                                                                       600
ccnnanaatn g
      <210> 358
      <211> 619
      <212> DNA
      <213> Homo sapiens
      <220>
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<221> misc_feature
       <222> (1) ... (619)
       <223> n = A,T,C or G
       <400> 358
ggtacttttt tttttttt ttttttttt ttgagatgga gtctcgctct gtcgcccagg
                                                                              60
ctggagtgca gtggcgcaat ctctgctcac tgcaacctcc gcctcctggg ttcaagcaat
                                                                             120
tetectgtet cageeteeca aatagetggg attaegggea tgtgteaega egeteggeta
                                                                             180
atttttgtat ttttagtcga gacgaggttc caccatgttg gctaggctgg tctcaaactc
                                                                             240
ctgacctcag gtgatccgcc tgcctcggcc tcccaaagtg ttaggattac gggtgtgagc
                                                                             300
cactgogoco agcaagcaac ctagatttta aaacaacatg agataaataa gootaattgg
                                                                             360
atttaactac atctaacatt tttactaata gttgnaatac tggtagaatt tggaaactat
                                                                             420
tatatatatt atgengaaaa gtaaataatt etggtaaaat canttanggn centgaattt
                                                                             480
nagcataggg gaaaaaaaga tgccntttta aatccaataa gtaaaaaccn tttaaccctn
                                                                             540
tntttaaatt ggaanttccc cccaatttnt tattaatttc aacttntttt gaaaactcat
                                                                             600
ntttccnaaa antnggggg
                                                                             619
       <210> 359
      <211> 624
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (624)
      <223> n = A, T, C \text{ or } G
      <400> 359
ggactttttt ttttttttt tttttttt ttttttggag gaaaacccgg taatgatgtc
                                                                              60
ggggttgagg gataggagga gaatggggga taggtgtatg aacatgaggg tgctttctcg
                                                                             120
tgtgaatgag ggttttatgt tgttaatgtg gtgggtgagt gagccccatt gtgttgtggt
aaatatgtag agggagtata gggctgtgac tagtatgttg agtcctgtaa gtagganagt
                                                                             180
                                                                             240
gatatttgat caggagaacg tggttactag cacagagagt tctcccagta ggttaatagt
                                                                             300
ggggggtaag gcgaggttag cgaggcttgt tanaagtcat caaaaagcta ttagtgggag
                                                                             360
tagagtttga agtccttgag agaggattat gatgccactg ngaatgcntt cctaatttga
                                                                             420
gtttgctagg cagaatagtn atgaggatgt aaacccctng gccaattatt aaaaatgact
                                                                             480
genecegtga aaettnaggg ggtttggatt aaaaangett gtaetteeaa nggetnintg geetnattta aaaaatttee etnnnenaat ttagggettn ttnnennaag eenanagggn
                                                                             540
                                                                             600
ccccnancct ttcccggggg ggcn
                                                                             624
      <210> 360
      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(611)
      <223> n = A,T,C or G
      <400> 360
acgcggggag gcggaggett gggtgcgttc aagattcaac ttcacccgta acccaccgcc
atggccgagg aaggcattgc tgctggaggt gtaatggacg ttaatactgc tttacaagag
                                                                             120
```

```
gttctgaaga ctgtcctcat ccacgatggc ctagcacgtg gaattcgcga agctgccaaa
                                                                         180
geettagaca agegeeaage ceatettigt gigettgeat ceaacigiga igageetatg
                                                                         240
tatgtcaagt tggtggaggc cctttgtgct gaacaccaaa tcaacctaat taaggttgat
                                                                         300
gacaacaaga aactaggaga atgggtaggc ctttgtaaaa ttgacagaga ggggaaaccc
                                                                         360
cgtaaagtgg ttggttgcag ttgtgtagta attaangact atggcaagga gtctcagcca
                                                                        420
aggatgtcat tgaagagtat ttcaaatgcc agaaatgaag aaattaaatc nttggcttac
                                                                        480
ttaaaaaaaa annnnnnnn aaaaaaaagg tccttgggcg gnacaccctt aaggggnaat
                                                                        540
tennnnecet gggggeentt ataangggnn cenaettggg ceaaattggg naaananggg
                                                                        600
naaanttttt n
                                                                        611
      <210> 361
      <211> 404
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (404)
      <223> n = A, T, C \text{ or } G
      <400> 361
acatatttta atagaaagat acaacctttt tattttcact ccttttattt ctgctgcttg
                                                                         60
gcacattttt gagttttccc acattatttg tctccatgat accactcaag cagtgtgctg
                                                                        120
gacctaaaat actgacttta gttagtatcc ttggattttt agattcccag tgtctaattc
                                                                        180
cctgttataa tttgcgcaaa caaaacaaaa tgttatgata atctttctcc actgttctaa
                                                                        240
tatatattgt attittattt gatagettgg gatttaaaac atctetgttg aaggettttg
                                                                        300
atccttttga gaaataaaga tctgaaagaa atggcataat cttaaaactt gataaaaaaa
                                                                        360
aaanannnnn nnnaaaaaaa aaagtacctn ggccgngacc acgc
                                                                        404
      <210> 362
      <211> 322
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (322)
      <223> n = A, T, C \text{ or } G
      <400> 362
ggtacttttt ttttttttt tttttttt ttttttggag ttgtaggcaa atgtttaatt
                                                                         60
aattetgete atatgeacat etgaaageat gagacaeact ceacagacag caegeactgg
                                                                        120
ggctggtggg gcanatgggc actcgccgat taggtattaa tgtcaataat acgtgcataa
                                                                        180
agtgctgata aaataactta agtgttacaa aaagagacag tccacggtgg ctgcaggcac
                                                                        240
atgcaggcgg gactgggtca gacactccag ggctgcacat gttccagctg gcctgagtcc
                                                                        300
gacacgtcat agctggcctt gt
                                                                        322
      <210> 363
      <211> 616
      <212> DNA
      <213> Homo sapiens
      <220>
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<221> misc feature
       <222> (1) ... (616)
       <223> n = A,T,C or G
       <400> 363
 cgaggtacgc gggctaagca agggaaaaat aacagtttct ctgagccaga gaagacttga
 tcacagttct ccaagcatcg tgatagcaat gcttaacccc aggaagattt caaggcaggg
                                                                                60
 agaagaacat ttcaaataag attcttgtta acccatttat gcctagtgtt ccattattgg
                                                                               120
                                                                               180
 aatgctaage ttgtgggagt catttacate ctactgetea aagteattge caaggtetga
                                                                               240
tttttcacac aaaaaattgc aacccccagc ataaatgttt agctactgtc atcagttagc
                                                                               300
aaattcatcc acacaaacac aattagagtt tggtttttt ttaagctttt caaaacttac
taaactggca caattttata tgtatgctat ttggtgnatt tatgcttaag agcnaaaaag tttgatggga ttttaaattc angccaagcc tacacgctga gacaatccct acaaccatgg
                                                                              360
                                                                              420
                                                                              480
nagtaactaa ngaaccttta tctaagnttt taagttttaa anggagngct taatggttca
                                                                              540
ngtctangtt ggaatttcct tcanaaattt cntcttttaa aaaattttcc caaaatnggt
                                                                              600
ccttaaaaaa ctcann
                                                                              616
       <210> 364
       <211> 618
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (618)
       <223> n = A, T, C or G
       <400> 364
cgaggtacgc ggggcttctc gcctaacgcc gccaacatgg tgttcaggcg cttcgtggag
gttggccggg tggcctatgt ctcctttgga cctcatgccg gaaaattggt cgcgattgta gatgttattg atcagaacag ggctttggtc gatggacctt acactcaagt gaggagacag gccatgcctt tcaaatgcat gcagctcact gatttcatcc tcaagtttcc gcacagtgcc
                                                                               60
                                                                              120
                                                                              180
caccagaagt atgteegaca ageetggeag aaggeagaca teaatacaaa atgggeagee
                                                                              240
                                                                              300
360
cgttttaaag ttatgaaggc aaagaaaatg aggaacagaa taatcaagaa tgaaagttaa
agaaacttca aaaggcagct nttctgaaag cttnttccca aaaaagcacc tgggtacctg
                                                                              420
geegggeegg eegtttaaaa gggenaatte caccaetgge ggeegtetan ngggateeaa
                                                                              480
                                                                              540
cttnggacca acttggngga atatggcnaa attgttcctg gggnaaatgt ttncgttcaa
                                                                              600
attncncaaa ttacggcc
                                                                              618
       <210> 365
       <211> 601
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (601)
      \langle 223 \rangle n = A,T,C or G
      <400> 365
acgtectgga ggactetatt gtggacceae agaateagae catgactaee tteacetgga
                                                                               60
acatcaacca cgcccggctg atggtggtgg aggaacgatg tgtttactgt gtgaactctg
                                                                              120
```

```
acaacagtgg ctggactgaa atccgccggg aagcctgggt ctcctctagc ttatttqqtq
                                                                           180
tctccagagc tgtccaggaa tttggtcttg cccggttcaa aagcaacgtg accaagacta
                                                                           240
tgaagggttt tgaatatatc ttggctaagc tgcaaggcga ggccccttcc aaaacacttg
                                                                           300
ttgagacage caaggaagee aaggagaagg caaaggagae ggeactggea getacagaga agecaaggae etegecagea aggeggeeae caagaacage ageageagea acagtttgtg
                                                                           360
                                                                           420
taaccagnet accaacaaca nagnacecca nacaggtagg ettacecett tggeeteett
                                                                           480
taatggacct tggccgggaa caccettang gcgaattcag ncactggggg ccgtactang
                                                                           540
ggatcenett ggaccaactt ggggaaacag ggcaaaattg ttettgggga aattntatee
                                                                           600
                                                                           601
       <210> 366
       <211> 321
       <212> DNA
      <213> Homo sapiens
      <400> 366
acttttttt ttttttttt tttttttgag atggagtctc actctgtcgc ccaggctgga
                                                                            60
atgcagtggt gcaatctcag ctcactgcaa cttccacctc ccaggttcaa gtgattctcc
                                                                           120
tgecteagee teccacatat etgggactae aggtgeacae caccatgeee agetaattte
                                                                           180
tttgtatttt ttagtagaga cggggtttca tcttattggg caggctggtc tcgaactcct
                                                                           240
aaccttgtga tetgeecace teggeettee aaagtgetgg gattacagge gtgagecace
                                                                          300
gtgctcggcc acccgcgtac c
                                                                          321
      <210> 367
      <211> 264
      <212> DNA
      <213> Homo sapiens
      <400> 367
actgatcatg gagttaatca acaatgtcgc caaagcccat ggtggttact ctgtgtttgc
                                                                           60
tggtgttggt gagaggaccc gtgaaggcaa tgatttatac catgaaatga ttgaatctgg
                                                                          120
tgttatcaac ttaaaagatg ccacctctaa ggtagcgctg gtatatggtc aaatgaatga
                                                                          180
accacctggt getegtgeee gggtagetet gaetgggetg actgtggetg aatactteag
                                                                          240
agaccaagaa ggtcaagatg tacc
                                                                          264
      <210> 368
      <211> 488
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(488)
      \langle 223 \rangle n = A,T,C or G
      <400> 368
ggtacagatg cacaggaggc catagggttt aggcanaggg gagcacaaan gttgaagatg
                                                                           60
aggegetgee ateaatgetg ggaetteagg enaagggeag gaactgagga ageeacaagg
                                                                          120
gaggacattt tetgeagttg etgaancagt ancaactagg teetgagaaa geeetntete
                                                                          180
gtggaagaat aacagccagg cnggaaagct tttcatcctg caaagctggg gaagaagatt
                                                                          240
cttccttaaa ttgtcatctg cacttcagct cangaatcct gttggctgaa gtccagagtg
                                                                          300
tccntttctg attcctgaag tanatnaaca gcccngnccc aangaagagn aggnntagta
                                                                          360
caaagcenne thegegtace tgthegggeg gnngttegna aggnteaaat teeagcacaa
                                                                          420
```

```
480
ttgnctgccg ttantagttg gattctnact ttngtactta ncttgqcqta ntttatgqtn
                                                                                 488
ataanttq
       <210> 369
       <211> 602
       <212> DNA
       <213> Homo sapiens
       <221> misc_feature
       <222> (1) ... (602)
       <223> n = A, T, C \text{ or } G
       <400> 369
acgggggttt cactacttct cccccggact ccttggtagt ctgttagtgg gagatccttg
                                                                                  60
ttgccgtccc ttcgcctcct tcaccgccgc agaccccttc aagttctagt catgcgtgag
                                                                                 120
tgcatctcca tccacgttgg ccaggctggt gtccagattg gcaatgcctg ctgggagctc tactgcctgg aacacggcat ccagcccgat ggccagatgc caagtgacaa gaccattggg ggaggagatg attccttcaa caccttcttc agtgaaacgg gtgctggcaa gcatgtgccc
                                                                                 180
                                                                                 240
                                                                                 300
cgggcagtgt ttgtagactt ggaacccaca gtcattgatg aagttcgcac tggcacttac
                                                                                 360
eggeagetet teaccetgag caacteatea eaggenagga aaaatgetge aataactate
                                                                                 420
ccgaaggcac tacaccattg gcaaggagaa taattgacct gtgttggacc gaattcgcaa
                                                                                 480
gctggctgac catgcaccgg cttaagggtt nttggttttc ccaacttttg gggggggaac
                                                                                 540
tgggtttngg gtaaccctnn tggtnatngg aacgntttta antggatttt gggaanaaan
                                                                                 600
                                                                                 602
       <210> 370
       <211> 257
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(257)
<223> n = A,T,C or G
       <400> 370
acttttttt tttttttt tttagttttt ttttattttt tacaaatata ctggagaatc
                                                                                  60
atgeaatget gecageattg gatgeaatee ggggeeacaa gtetgeacae teettigeta
                                                                                 120
etggteetgt aatggeagaa cettteatet egeetttatt gnteactatg acteetgeat
                                                                                 180
tatetteaaa ataaagaaac aegeeatett ttetaeggta tgaetttegt tgtegaatga
                                                                                 240
ccactgctgg atgtacc
                                                                                 257
       <210> 371
       <211> 607
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (607)
       <223> n = A, T, C \text{ or } G
```

```
<400> 371
 actititit tittitit tittitigct attiagtitt tatticataa tcataaacti
                                                                              60
 aactctgcaa tccagctagg catgggaggg aacaaggaaa acatggaacc caaagggaac
                                                                             120
 tgcagcgaga gcacaaagat tctaggatac tgcgagcaaa tggggtggag gggtgctctc
                                                                             180
 ctgagctaca gaaggaatga tctggtggtt aagataaaac acaagtcaaa cttattcgag
                                                                             240
ttgtccacag tcagcaatgg tgatcttctt gctggtcttg ccattcctgg acccaaaggg ctccatggcc tccacaatat tcatgccttc tttcactttg ccaaacacca catgcttgcc
                                                                             300
                                                                             360
atccaaccac tcagtcttgg caagtgcaga tgaaaaactg ggaaccantt ggggttgggt
                                                                             420
ccacatttgc catggacaag aatgccagga acccgtatgc tttaaggatg aagtctcatc
                                                                             480
ttcaaaattc ttccccataa atggacttgc caccagngcc attatggcgt gtgaagtccc
                                                                             540
cancetggee cataaaccet ggaaaaatnt tggnaaaccg gaaccetttt aaccaatcet
                                                                             600
ttttttc
                                                                             607
       <210> 372
       <211> 607
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (607)
       <223> n = A, T, C \text{ or } G
       <400> 372
acgaatgtgg gaattactca ggagcagcag aatatettta tttttttaga gtgetggtte
                                                                              60
cagcaacaga tagaaatgct ttaagttcac tetggggaaa getggeetet gaaatettaa
                                                                             120
tgcagaattg ggatgcagcc atggaagacc ttacacggtt aaaagagacc atagataata
                                                                             180
attetgtgag ttetecaett cagtetette ageagagaac atggeteatt caetggtete
                                                                             240
tgtttgtttt cttcaatcac cccaaaggtc gcgataatat tattgacctc ttcctttatc agccacaata tcttaatgca attcagacaa tgtgtccaca cattcttcgc tatttgacta
                                                                             300
                                                                             360
cagcagtcat aacaaacaag gatgttcgaa aacgtcggca ggttctaaaa agatctaggt
                                                                             420
taaaggttat tcaacangga gtcttacnca tntaagaccc cattacngga atttggtgaa
                                                                             480
tggttatatg taactttgac tttaangggc tcaaaaaaag ctnaggggat gtgaatcaag
                                                                             540
cttgngaagg ctttttttgg gggctngntt nngggtttnt tgnaaagncc ngttttnntt
                                                                             600
ttggaat
                                                                             607
       <210> 373
       <211> 618
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (618)
      <223> n = A,T,C or G
      <400> 373
acttttaatg tttgctgttc aaacgaaaat agattggatc ttggttaagt tcacttggtt
                                                                              60
tggccaggca cagtggctca cgcctgcagt cccagcactt ggggaggtgg aggcgggccg
                                                                            120
atcacctgag gtcaagagtt tgagaccagc ctggctaacg cggtgaaacc ccatttctac
                                                                            180
taaaaataca aaaaattagc tgggcgtggt ggtgcgcgct tgtaatccca gctactcggg
                                                                            240
aggctgaggc aggagaatcg cttgagccag agaggcaaag gttgcaataa gccaagatag
                                                                            300
cgccattgta ttccagcttg gacaacaaga gcgaaactct gtctaaaaaa aaaaaaaaa
                                                                            360
```

```
cacacacaca acacaatatt ttcacgcctg taaacctagc acattgggaa gccaaggtgg
                                                                         420
gaggattgct tgaggccagg agttcaaggc ttgcantgag ctatgaatgn acactgnacc
                                                                         480
tttggnegng aacacnetta nggccaaatt cengcacaet tgggggeegg tactaanggg
                                                                         540
atcccanctt tggnnccaaa nttggngnaa acatgggcaa aattggtncc tggngaaaat
                                                                         600
ggttccgttc caaatccc
                                                                         618
       <210> 374
       <211> 605
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(605)
       \langle 223 \rangle n = A,T,C or G
       <400> 374
acccagctgc tgcccacatt tetggtccag agtcccgaac cccgagcact gggatgcctg
                                                                         60
gctactccga gcgttatcca gactagcgag tgggaggcag atgtaaaatc tggaacgcag
                                                                        120
attttagttt gttggaagga gaaatgtaac atagtgaacc acgcatctct ggagggtgta
                                                                        180
aagcagagac agccaagagc caaggcactg atgtttgaac tggaaacttc aaaacgttta
                                                                        240
ataagagtet teaggatggg tttgaactag acaagetaga aatttettta gaacaccage
                                                                        300
tctagcatgc atctcccact tttggctttc ctggagagga gcttgaagag gtggttctgc
                                                                        360
agacagecae agtgataete aggaaacnea gaggaatgga tttgaetttt etgetaggaa
tetttggtat aagtteteet tgagttgtaa gangeatgga aatatacatg aaactgaana
                                                                        420
                                                                        480
acctgcaagg aanggaaatg ggaacntttc atctgagtgn aaactaacca agtnggcaat
                                                                        540
ttngacttga aaccettgaa accttcnagt ccaantectg gtttggggga taaangaacc
                                                                        600
ggncn
                                                                        605
      <210> 375
      <211> 602
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(602)
      <223> n = A, T, C \text{ or } G
      <400> 375
acggatgcta cttgtccaat gatggtaaaa gggtagctta ctggttgtcc tccgattcag
                                                                         60
gttagaatga ggaggtetge ggetaggagt caataaagtg attggettag tgggegaaat
                                                                        120
attatgettt gitgittgga tatatggagg atggggatta ttgctaggat gaggatggat
                                                                        180
agtaataggg caaggacgc tectagettg ttagggacgg ateggagaat tgtgtaggeg
                                                                        240
aataggaaat atcattcggg cttgatgtgg ggaggggtgt ttaaggggtt ggctagggta
                                                                        300
taattgtctg ggtcgcctag gaggtctggt gagaatagtg ttaatgtcat taaggagaga
                                                                        360
atgaanagaa gtaagccgag ggcgtctttg attgtgtagt aagggtggaa ggtgatttta
toggaatggg aagtgattno taaggggntg tttgancocc gtttgtgcca gaatangaag
                                                                        420
                                                                        480
tggaatgett ettanggett caataaatga anggeanaat gaattgaaag gtaaanaaac
                                                                        540
cntnaagggt ggacttgtta ctgataaccn tcctaaaatc attgccccgn aacttggccg
                                                                        600
gg
                                                                        602
```

<210> 376

```
<211> 611
       <212> DNA
       <213> Homo sapiens
       <221> misc_feature
       <222> (1) ... (611)
       <223> n = A,T,C or G
       <400> 376
 acgegggate gaagaattea caaaaaacaa tageeteate ateeceacea teatageeac
                                                                          60
 catcaccctc cttaacctct acttctacct acgcctaatc tactccacct caatcacact
                                                                         120
actececata tetaacaaeg taaaaataaa atgacagttt gaacatacaa aaceeacee
                                                                         180
attectecce acacteateg coettaceae getactecta ectatetece ettttatact
                                                                         240
300
aaaaaaaang tncngccatt tttngtttcn ggtaaacngg aatataangn gaaagaacaa
                                                                         360
acnttggaac atacttaatg gatttttata gaactttgna aaccaaagga gattcatgtt ttanaagtct ggccttttt atatcttgga agaaaattat gtntggaggc tntaaataaa
                                                                         420
                                                                         480
teccattatt ttetcaggga atetgggtag gaattgeegg catgggaant tttnngggge
                                                                         540
cggatnggaa agtttggcct aanaaatngc nctttntnaa naattttgga attttgggaa
                                                                        600
gcccnaagca n
                                                                        611
      <210> 377
      <211> 367
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(367)
      <223> n = A, T, C or G
      <400> 377
acgcgggccg tttggcatct ctgccctcat cgtgggtttc gactttgatg tcactcctag
                                                                         60
gctctatcag actgacccct cgggcacata ccatgcctgg aaggccaatg ccataggccg
                                                                        120
gggtgccaag tcagtgcgtg agttcctgga gaagaactat actgacgaag ccattgaaac
                                                                        180
agatgatetg accattaage tggtgateaa ggcaeteetg gaagtggtte agteaggtgg
                                                                        240
caaaaacatt gaacttgctg tcatgaggcg agatcaatcc ctcaagattt taaatcctga
                                                                        300
agaaattgag aagtatgttg caaaaaaaa aananaaatn aaanaagtac ctcggccgng
                                                                        360
accacgc
                                                                        367
      <210> 378
      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(611)
      <223> n = A, T, C \text{ or } G
ggtacctgga tcctgctcct ctgggttgaa acccgggcgc cgccaagatg ccggcttacc
                                                                         60
```

....

```
actettetet catggateet gataccaaae teateggaaa catggeactg ttgeetatea
                                                                             120
gaagtcaatt caaaggacct gcccccagag agacaaaaga tacagatatt gtggatgaag
                                                                             180
ccatctatta cttcaaggcc aatgtcttct tcaaaaacta tgaaattaag aatgaagctg
                                                                             240
ataggacett gatatatata actetetaca tttetgaatg tetgaagaaa etgeaaaagt geaatteeaa aageeaaggt gagaaagaaa tgtataeget gggaateaet aatttteeat
                                                                             300
                                                                             360
tectggagag cetggtttte caettaacge aatttatgee aaacetgeaa acaaacaggg
                                                                             420
aagatgaagt gatgagagcc tatttacaac agcttaaggg caagaaactg gactggaact
                                                                             480
ttgtgaagaa gttttcgacc cttagaatgg ttaaaccnac agtgggggga cttgcttttg
                                                                             540
gaaaanaccg tttattgacn anagtttttt tggactggan atgaaaggng cccnggttng
                                                                             600
ccccaatttn n
                                                                             611
       <210> 379
       <211> 602
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(602)
       <223> n = A, T, C \text{ or } G
       <400> 379
acagetggtt ggacetatte atgeatette accageaget ggageatete caccettggt
                                                                              60
atttetggtg taaattaett gagetetgtg etttgaaace agtttgataa gteetttaet
                                                                             120
aaggagetee tgaagggetg eeetggeeag ggageetega atetteagte teteagagae
cacagetggg gttataagtt tatagttggg aactteetta cagagtttat cataggtage tttgtcaaac aagactaagt tattgagett gtcccgaact ttgcctttgg accacttett
                                                                             180
                                                                             240
                                                                             300
ctttttggcc ttgcccccgg atttgttcac tgggtctttg nctttcttgg ccgactttcc
                                                                             360
agegteette ttettettgt egteettagg eggeattgee aageteggag aatageanea
                                                                             420
gacacngnaa cctngtcaag atgtcngaca aaaagccccg ggtaccttgg gcgngaacac
                                                                             480
gcttaaggcg aattecacae actggcggcc gtactanggg gatccagett nggaccaact
                                                                             540
tggnggaaac atggcnaact gnttcctngn ggaaaatgtn atccgttaaa attnccccaa
                                                                             600
                                                                             602
       <210> 380
       <211> 598
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (598)
       <223> n = A, T, C or G
       <400> 380
ggtacngcgg ggggtgcctg gctccgtttc ctgcttttgg ttcttacagt agtcggcgta
                                                                             60
ggccttagat tttttactgt ctcctgaaga atttaacaca aacatggata tcagaccaaa
                                                                            120
tcatacaatt tatatcaaca atatgaatga caaaattaaa aaggaagaat tgaagagatc
                                                                            180
cctatatgcc ctgttttctc agtttggtca tgtggtggac attgtggctt taaagaccat
                                                                            240
gaagatgagg gggcaggcct tigtcatatt taaggaactg ggctcatcca caaatgcctt
                                                                            300
gagacageta caaggattte cattttatgg taaaccaatg cgaatcagta tgcaaaacag
                                                                            360
attecggata taatateaaa aatgegtgga aettttgttg ecaagaaaag aanaaagaaa
                                                                            420
agaaaaagnc caaacttggg aacaactgna caaccncaac caaaaanctg ggcnngggac
                                                                            480
```

```
tecaaateae ttataeeeag ggaatteaee eenaatetta ggteetgata eetteaaeta
                                                                      540
tatttaatcc ttaaaactta nccgaagagc taatngatga tgtntcctgc cggtaacn
                                                                      598
      <210> 381
      <211> 631
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (631)
      <223> n = A, T, C \text{ or } G
      <400> 381
ggtacgcggg gagagtgtgg tcaggcggct cggactgagc aggactttcc ttatcccagt
                                                                      60
tgattgtgca gaatacactg cctgtcgctt gtcttctatt caccatggct tcttctgata
                                                                      120
180
ctcggtcaaa agaatctgtt ccagaattcc ccctttcccc tccaaagaag aaggatcttt
                                                                      240
ccctggagga aattcagaag aaattagaag ctgcagaaga aagacgcaag tcccatgaag ctgaggtett gaagcagetg gctgagaaac gagagcacga gaaagaagtg ctttagaagg
                                                                      300
                                                                      360
caatagaaga agaaccacaa cttcgtaaaa atggcngaan aagaaacega ccnccaaaat
                                                                      420
gggagettat taaagagaan ceagangnne caatngnttg geeaactggg accgtttgea
                                                                      480
anaagaaggg ttagccccnt tgaanaaatg ccggaagaac caaagaattc caagaccctt
                                                                      540
gntgenaaac ttgaacttge ctaattggte ttgagaactg ettttteee atceetteta
                                                                     600
aaatccaaaa atgnacctgc ccgggggccg t
                                                                     631
      <210> 382
      <211> 613
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(613)
      <223> n = A, T, C \text{ or } G
      <400> 382
60
aaagettttt gaaactatgt tttctccagg gaagttctct ttcaacaaga tggttttcac
                                                                     120
tactgataac ttaacatgct ggaaacctgg taatgtttct atgactttat tttctaacat
                                                                     180
cttctttaaa tctttaggca tagcatgctc tttggcagct ctcaaggagg gctgtttcca
                                                                     240
tgtggctcca agttccttga actgctggct gcactgagtg gactgtctgt gtcttgagag
                                                                     300
ggagetgeat titeattgae ttatggteec acaagtgace etgaggeaan gtenaattgg
                                                                     360
tetneanaac atttttggee etetettete etttttgaet tttetgagae tgaeagttet
                                                                     420
tttganggaa tccagggnna angetteent etetaatggg ggntaaatte attttccaaa
                                                                     480
anggneggtt tttgggaaaa tnaaanttga aanggeatee nttttattaa tgeecenane
                                                                     540
ttttaanttc ngattntnaa cttnctgnta gaatttgtgg atccnccaaa ttggcttaat
                                                                     600
attcaaatag ctt
                                                                     613
      <210> 383
      <211> 628
      <212> DNA
      <213> Homo sapiens
```

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<220>
       <221> misc feature
       <222> (1) ... (628)
       <223> n = A, T, C or G
       <400> 383
ggtactttga ccctggaaag gtatgggtct gcttaaaaga aagaagaaac atacacgtaa
tcaaataaag cttaacatta tgcagggctt ataatcattt tcagcaacgg actgcaagct
                                                                              120
gcactgtgaa gaaaatgcat agcagaggag aaagctgggg atctgaggaa ataggtaagg
                                                                              180
aaaacagtgt caacacacag tggaagaagt gatgaagaca tctattccgg agctcacgtg
                                                                              240
ccatgecetg ctagegttee ttaacaagee acctgeteea gaaggeeaca geetgaeeet
                                                                              300
cccaagtgga atataaatgc ccaagtgcca catgaagcca ccttctncac tacctaaaaa
                                                                              360
ggttgtctgg gactgagctc agaacacaca cctttctggg ctaccaaacc tttaagtgga aagaattttt tnctaaatat ctanttttna tacccacttt aacgccactt ttatattgaa
                                                                              420
                                                                              480
attgggette taattagnee ettteeteaa tteettagga nggaacteat aatgggagee
                                                                              540
aaccaaccag ggattctacc cccaatngac tgnnctttaa angtattatt aattttgang
                                                                              600
ggcaaaggtg tgaatggttt acaatacc
                                                                              628
       <210> 384
      <211> 620
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (620)
      <223> n = A, T, C or G
      <400> 384
acaggtaage eetggetgee tecacecaet eccagggaga ecaaaageet teatacatet
                                                                               60
caagttgggg gacaaaaaa gggggaaggg ggggcacgaa ggctcatcat tcaaaataaa acaaaataaa aaagtattaa agcgaagatt aaaaaaaattt tgcattacat aatttacacg
                                                                              120
                                                                              180
aaagcaatge tateacetne eetgtgtgga ettgggagag gaetgggeea tteteettag
                                                                              240
agagaagtgg ggnggctttt angatggcaa gggacttcct gtaacaatgc atctcatatt
                                                                              300
ttggaatgac tattaaaaaa acaacaatgt gcaatcnaaa gtctcggccc atttgcggaa
                                                                              360
ctttgggggg atgettgett enacegantt ggtgncaace tttnnceggt tecanttttt
                                                                              420
naaattetta gtnnaagenn aaaaanntag aatanenena nancataaet tannaaneea
                                                                              480
tttaanaggt ccctcggccg gaacnnnctt aanggtnaat cccantnnnt ggcgggcgtt
                                                                              540
actnenggat ccancettgg nnecaaantn gnggaattea tggennaace gnteetgggn
                                                                              600
gaantngttn ccttnaaanc
                                                                              620
      <210> 385
      <211> 535
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (535)
      \langle 223 \rangle n = A,T,C or G
      <400> 385
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```
ggtacttttt tttttttt tttttttggt atttagtttt tatttcataa tcataaactt
                                                                        60
aactetgeaa teeagetagg catgggaggg aacaaggaaa acatggaace caaagggaac
                                                                       120
tgcagcgaga gcacaaagat tctaggatac tgcgagcaaa tggggtggag gggtgctctc
                                                                       180
ctgagctaca gaaggaatga tctggtggtt aagataaaac acaagtcaaa cttattcgag
                                                                       240
ttgtccacag tcagcaatgg tgatcttctt gctggtcttg ccattcctgg acccaaagcg
                                                                       300
ctccatggcc tcacaatatt catgccttct ttcactttgc caaacaccac atgcttgcca
                                                                       360
tccaaccact cagtettggc agtgcagatg aaaaactggg aanentttgg ggtngggnen
                                                                       420
acatttgeet tggccaaaat geenggaace ggeeeegtae ettgneengg eeggeeggtt
                                                                       480
caaaagggcg aattccacac acttggcggg ccgtactang gggatccaac ttcgg
                                                                       535
      <210> 386
      <211> 642
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(642)
      <223> n = A,T,C or G
      <400> 386
acagcattgg cagtggtgcg tcagaggtgg cagaactatt tcacactaac cagttgaaga
                                                                        60
ctacacaaga ttaataccat ccagcatcag gatatagctg tggattttac aaaccattct
                                                                       120
tatttctaac ttcaggagtt gatgtttttc ccagtccatc ttaaaatatt actgctttaa
                                                                       180
tcacagatca ggtaaaaagg acaacatgca caacctccac ctagaatcct gttgtagcct
                                                                       240
agacagtgaa atgatatgac atcagaagac tttaaaaattg cagctccttt tggatccccc
                                                                       300
aaagtgtatc tgcactcttc ttcaaacggg ccctctttcc tcaagaagtc agaagtcacc
                                                                       360
ttcacaangn ctgagaattc cattctgnnc ccaaantgca agggacactn aaggaagaca
                                                                       420
teattettt atteegtnaa agaecettaa tteatgggng gaaaetgggt geaceegeet
                                                                       480
nagaatettt attanaetet ttgnecaatt tggttacaga agagntnean taneeceang
                                                                       540
aannggtage etttggagtt tgantcacce tcataagcac eettaaacca eetgnttggg
                                                                       600
gaaccttctt tcactggtcc ctaactttat tangccctaa ag
                                                                       642
      <210> 387
      <211> 256
      <212> DNA
      <213> Homo sapiens
      <400> 387
ggaccttttt ttttttttt tttttttt tgaaaagaaa ggccttacat atttattact
                                                                        60
gaatccagec aaccaacgtg ttcataacag attcagagag gaaaacacgt cgaaatctcc
                                                                       120
agatagtggt gacattttca gcttgatatg gtaacatgat cgtgaccttc agacagcata
                                                                       180
aatatgtgtg ccatctcatg tgcaattcct tatagaccca gcttggttct tctccaatgt
                                                                       240
ctccttttgg agttgt
                                                                       256
     <210> 388
     <211> 566
     <212> DNA
     <213> Homo sapiens
     <220>
     <221> misc_feature
     <222> (1) ... (566)
```

PCT/US99/19424 WO 00/12702

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120

180

240 300

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agaactacaa aggcaggagg tgtaagtgaa tttttattgg gaggggaggt tggcaactta

aacagcagca aataaagagt gaataaggaa actccctgtt gccacagata cacaagacct ccgtatgtga tacaggagcc atttcaattt gtgaccccta gacagagatg gcaagtgctt

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                                                                           480
ttatgtggaa tetettaaat tgnaattact acatttetta atttecaggt atnecaaaca
                                                                           540
cagtconttg caaaactggt cagntactta aatnoongat coattttagg onttacataa
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tggtggaagc agcaggggag gatgagcggg agctggccgc agagatggca gcagcattcc
                                                                           180
tcaatgaaaa cctccctgaa tccatctttg gagctcccaa ggctggcaat gggcagtggg
                                                                           240
cctctgtgat ccgaqtgatg aatcccattc aagggaacac actggacctt gtccagctgg
                                                                           300
                                                                           360
aacanaatga ggcagnttta gtgtggctgt gtgcaaggtt tccacactgg tgaagactgg
tntgtgctgg tgggtgtngn canaggacct ngntnctaaa accncgnntt tgggcaatgg
                                                                           420
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                                                                           480
                                                                           540
ttnatatngt natggnnaaa cntntanccg nnntntaatc ttggaatata tatnaatacc
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tecagatgee gtgaatttaa etattegtta eaggettgte etgeaatatg etetggagea
                                                                           180
acttgcctqc agagatttct qtatccacgg cttcagagca gaaagagaaa gcaaagaagt
                                                                           240
                                                                           266
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120
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aaaaaaaata attacaggtt agatatttaa tccaaggttt aacatgggga tgatctcata
                                                                       180
aggcaatttc tttcctttaa taaatattaa agtgaatatt attctggaag caaatcatct
                                                                       240
cctaattett catcagcaaa atcateetea tegateettt tettggetge agtttttggt
                                                                       300
cgttctattt gagggccaag tgggtccaca taggaggcat ctatttcttt gntactgcta
                                                                       360
ctttcataag gntcatttgt cccaggtaaa agctctgagt ctggccttan tccgtcaccc
                                                                       420
                                                                       480
tttactactq genetatagt etggecacta tnaacgntag cettnettnt enttttgnea
enggageece caatgeannt tingentgae titageneng gneectaatt etteatitit
                                                                       540
                                                                       600
ccacctttna gnttttggca antcttgagc cntttttaat cnaagacttn gcanagccaa
                                                                       611
ttaaaaaccc c.
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                                                                        120
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aaaagaatgo caaaatcago agtotootgg aggagcagtt ccagcagggo aagcttottg
                                                                        180
cgtgcatcgc ttcaaggccg ggacagtgtg gccgagcaga tggctatgtg ctagagggca
                                                                        240
aagagttgga gttctatctt aggaaaatca aggcccgcaa aggcaaataa atccttgttt
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tgtcttcacg caaaaaaaaa aaaaaaaaaa aaaaagtacc
                                                                        340
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      <211> 557
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      <221> misc_feature
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                                                                         60
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                                                                        120
agtcaaggaa actcattcac acttcaggtc cttctcctcc aggaaccagc attgttatat
                                                                        180
tatttccatt tagcaaaatc tgatgtaatt tagtaatcct tettccttct ggtgtgattt
                                                                        240
caaactcagt gacatcttcc agtactttnt ttttttttt ttttttttgg gtgttgagct
                                                                        300
tggacgettt ettaattggt ggetgetttt aggeetacta tgggtgttaa atttttaete
                                                                        360
                                                                        420
tetetacaag gnttttteet agtggeeaaa agaagetggt eestettttg gaetacegtt
aaaattacca nggggattta aaangggtnt tgngggccaa attnaaagtt ngactangan
                                                                        480
totatttttg gocaaccaqt nttaaccaqq ottogqtanq qttqqccqcc cccgggtacc
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ttgggccggg aacacnc
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      <221> misc feature
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                                                                          120
                                                                          180
aaaggaaatc tgaaactcag caccgaggct ctgctcccca ctctgagagt gatctaccag
agcaggaaga ggagattctg ggatctgatg atgatgagca agaagatcct aatgattatt
                                                                          240
                                                                          300
gtaaaggagg ttatcatctt gtgaaaattg gagatctatt caatgggaga taccatgtga
tccqaaaqtt aggctgggga cacttttcaa cagtatggtt atcatgggat attcagggga
                                                                          360
agaaatttgt ggcaatgaaa gtagttaaaa gtgctgaaca ttacacttga aaccagccta
                                                                          420
gatgaaatcc ggttgcttga agtcagttcc aattcagacc ttatggatcc aaatngaaaa
                                                                          480
atggttgtca actactagat gactttaaaa ttcaggagtt aatggaacac atatttgcat
                                                                          540
gggatttgaa gttttggggc anattngtta agnggttctc aaatcaattn ttangggctt
                                                                          600
                                                                          617
tcctgccttg ggtnaaa
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      <221> misc feature
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                                                                          120
                                                                          180
coggtetece tgeccgteac cettggacag coggeeteca tetectgcag gtetggtgaa
actictettt acgaagatgg aagcacetac ttgagttggt ttcaccagag gccaggccaa tctccgaggc gcctgattta taaagtttct aaccgggact ctggggtccc agacagattc
                                                                          240
                                                                          300
ageggeagtg ggteaggeae ttattteaeg etgaaaatea acagggtaga ggetgatgat
                                                                          360
gttgggaatt attactgcat gccanggtca aactggcccg tcacttttcg gngaaggacn
                                                                          420
aaaggtggcc natcaaacca actgnggctt gaccattggc ttcatnttcc cgccatttga
                                                                          480
                                                                          540
taaccantga aatctggact gctttgtggg ngcctgctga aaacttntat nccnanaggc
cnaagtcatg acagtttttc natttactcg aaaaatntgg aaatgataat tttn
                                                                          594
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      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
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      <223> n = A,T,C or G
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acagtggtcc ttttcagagt tggacttcta qactcacctg ttctcactcc ctgttttaat
                                                                           60
                                                                          120
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gtctgtgcag tttctgacac ttgttgttga acatggctaa atacaatggg tatcgctgag
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actaagttgt agaaattaac aaatgtgctg cttggttaaa atggctacac tcatctgact
                                                                          240
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cattetttat tetattttag ttggtttgta tettgeetaa ggtgegtagt ceaactettg
                                                                          300
gtattaccct cctaatagtc atactagtag tcatactccc tggtgtagtg tattctctaa
                                                                          360
aagetttaaa tgtetgeatg cagecageat teaatagtga atggnetete tttggetgga
                                                                          420
attaccaaac tcagagaaat gnggcatcag gagaacatct taaccccatg aanggataaa agccccaaat ggngggnact tgataatagc nctaatgctt taaanatttg gtccactttt
                                                                          480
                                                                          540
tacctaaggt gagcccattg aaccannggt gctaaangct catacttcca actgaaatgg
                                                                           600
                                                                           611
ttaaggaaaa a
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                                                                           120
                                                                           180
tctgggactg tgaaacagga aagcagctgg cccttctcaa gaccaattcg gctgtccgga
                                                                           240
cetgeggttt tgactttggg ggcaacatca teatgttete caeggacaag canatggget
accagigett igigagetti titgacetge gggateegag ecagaitgae aacaaigage ectacaigaa gateeetige aatgacteta aaateaccag igetgiitigg ggaceeeting
                                                                           300
                                                                           360
 gggagtgcat catnetggce atgaaaagtg gagagetnaa ccagtattag tgeennagtt
                                                                           420
 tnnanaaggt gttngttnaa tgttaaagga gcantttccg gnagaataac cnacnttcag
                                                                           480
                                                                           540
 gttatteenn gganatgace anngtttnga eccettnnna gtecattaat neenaacttt
 tttachetca aattttnaan tnanaaaact tttngnatna aattnttnaa ttanttgtte
                                                                           600
                                                                           614
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 gettttteca aagateagge etaaagaaac agateacaaa agtettteat eateteetga
                                                                            180
 taaacgagag aagtttaaag aacagagaaa agcaacagtg aatgtgaaga aagacaaaga
 agataaaccc ttaaagacag aaaagcgacc caagcagcct gataaagaag gaaagttaat
                                                                            240
 ctgttctgaa aaggggaaag tgtcagagaa aagtcttccc aagaacgaga aggaagacaa
                                                                            300
                                                                            360
 ggaaaacatt tccgaaaatg acagagagta ttctggagat gcccaagtgg ataagaaacc
                                                                            420
 tgaaaatgac attgtgaaga gtccacaaga aaacttgagg ggaaccnaaa ngaaaacgag
 geagacecee ttecataget netactgetg gggattnaaa etttaaaett tggeacecat
                                                                            480
                                                                            540
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 gttgncaaaa ntttttncng ggaaagtcaa aaacttcttt gaaaaccttg ccnangattt
                                                                            600
                                                                            612
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                                                                                  120
teggacgeae aggatateca ttecatecae teteagecea ggaatgaaat egeetetett
                                                                                  180
gtagtaatca gtgctggctg ccgctctctc aacagacgtt cccattccat agcgattatt
                                                                                  240
ctcacagatg aaaatacaag gtaatttcca caaagctgcc atgttgtaag cttcgaatat ctggccctgg ttagcagcac catcgccata taaagtcagg cagacctcat cttttccatt
                                                                                  300
                                                                                  360
atacttacag gctagagcaa tcccagcgcc caagggcacc tgcgctccta cgatgccatg
                                                                                  420
gecegtana agtettggea tacatgtgea tegatectee tteetttage acaaneteet
                                                                                  480
tttgncctgt aactgcaaaa tttntcggac ggaaaggccc cggtgnaaag taaagccgtg
                                                                                  540
                                                                                  600
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                                                                                  601
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                                                                                  120
                                                                                  180
gaagacgacc ctgacaagcg catctcgatc tgctcctctg acaaacgaat tgcctgtgag
                                                                                  240
gaagagttet eegattetga agaggaggga gaggggggee geaagaacte tteeaactte
aaaaaagcca agagagtcaa aacagaggat gaaaaagaga aagacccaga ggagaagaaa
                                                                                  300
gaagtcaccg aagaggagaa aaccaaggag gagaagccag aagccaaagg ggtcaaggag gaggtcaagt tggcctgaat ggacctnttc agctctggct ttctgctgag tccctacgtt ctttcccaac cccttaaatt tataatttct attctctggg gatttatata aaaatttatt
                                                                                  360
                                                                                  420
                                                                                  480
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aatatgacca ctacttgatc ccaaacgccc tgctggagct ggccctgctg cttatggagc
                                                                        180
                                                                        240
aagacagaaa cgaagaggcc atcaaacttt tggaatctgc caagcaaaac tacaagaatt
                                                                        300
actccatgga gtcaaggaca cactttcgaa tccaggcagc cacactccaa gccaagtctt
ccctagagaa cagcagcaga tccatggtct catcagtgtc cttgtagctt tgtgcagcag
                                                                        360
                                                                        420
ttccgggctg gaagacagag acagctggac agagctcctg aaaacatttc aaaaataccc
                                                                        480
cetteceetg geetgeeetg cetttggggt ceaneggeae tteagttgga tggcacaace
tantgtatcc gtgcnnaaan cnaacctggc attttcaccc anntanccaa gggcttttgc
                                                                        540
caagggnana acagtggagc ccttggcttg ncctataaac atacgggtac cttggccgnn
                                                                        600
                                                                        604
acnn
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                                                                        120
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                                                                        180
ctgctgctgg tcattcatta atgtgtcaga catttaatca ggatgctgga ccttcaaaat
aactgaaaaa agaaccaaga aaaggcgttt ttgttttcaa caaactttac taaataaccc
                                                                        240
cggaaaggca atgaacgatc tgacaattta agctctaatg atttaaagct cagctagaag
                                                                        300
aaagtgaggc atgacatata ctgtcaacgg agggtgaagg aggcagattt ctggaaatgc
                                                                        360
aatgatccca cacatttgct tcaaggagaa acctgcagac atattttcag gtcttgctaa
                                                                        420
                                                                        480
gtaacaactg gttatttgta atcaatcatt tgggaaagtc tgctatgtag ctaanggcac
                                                                        540
tgtgaccccn gacaacngat gaaaaggaaa aagcnttgac agcaggaaaa atccttccat
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cttaaagaat ttaggggaca cctttaaagg aaaaaaattg ntccagcctc atttttacaa
                                                                        604
ntnt
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                                                                          60
ttccatgttt ttaaaagatt actttctact ttgtgtttca cagacattga atatattaaa
                                                                         120
ttattccata ttttctttc agtgaaaaat tttttaaatg gaagactgtt ctaaaatcac
                                                                         180
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```
240
ttttttccct aatccaattt ttagagtggc tagtagtttc ttcatttgaa attgtaagca
teeggteagt aagaatgeee atceagtttt etatatttea tagteaaage ettgaaagea
                                                                       300
totacaaato tottttttta ggttttgncc atagcatcag ttgatcotta ctaagttttc
                                                                       360
                                                                       420
atggggagac ttccttcatc acatcttatg ttgaaatcac tttctgtagt caaaggtata
                                                                       480
ccaaaaccaa tttatcttqa actaaattct aaagtatggg tatccaacca tatacatctg
                                                                       540
ggtaccaaac ataaatgctg acattentat attatagtna aggettaate nacttgcagg
tgaatggaaa aaaaataagc ttnaacctag gattctggaa tgaggaatgc tcn
                                                                       593
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      <221> misc feature
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gagtgcagtg gcgcaatctt ggctcactgc aacctctgcc tcctgggttc aagtggttct
                                                                       120
catgcctcag cctcctgggt agctgggatt acagacaagc accaccacaa ccagctagtt
                                                                       180
ttttttgttt tgttttttg agacggagtc tcgctctgtc accaggctgg agtgcagtgg
                                                                       240
                                                                       300
cacaatettg getcactgea acctetgeet cetgggttea agagattete etgetteage
ctnccaagta gctgggacta caggtgcaca ccatcacacc tggctaattt ttgtattttt
                                                                       360
aagtanagac ggggtttcac catgttggcc aggetggtct caaactcctg acctcaagtg
                                                                       420
aaccggccgc ttancctcca aagtgctggg attacaggcg tgagcccact ggcctggctg
                                                                       480
accatttggt tattaacagg gcccccaana tgcnccttta ngtgaaaggg natggcccca
                                                                       540
                                                                       591
gggaacaatt nngctgaaaa acaccaaagg conantccat aattonttgg n
      <210> 407
      <211> 463
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (463)
      <223> n = A, T, C or G
      <400> 407
                                                                        60
ggtactgatt ttaaaaaacta ataacttaaa actgccacac gcaaaaaaaga aaaccaaagt
                                                                       120
ggtccacaaa acattctcct ttccttctga aggttttacg atgcattgtt atcattaacc
                                                                       180
agtettttae taetaaaett aaatggeeaa tigaaaeaaa eagttetgag aeegttette
caccactgat taagagtggg gtggcaggta ttagggataa tattcattta gccttctgag
                                                                       240
                                                                       300
etttetggge agaettggtg acettgeeag etceageage ettettgtee actgetttga
                                                                       360
tgacacceae egeaactgte tgteteatat caegaacage aaagegacee aaaggtggat
agtotgagaa gototcaaca cacatgggot tgocaggaac catatcaaca atggcagcat
                                                                       420
                                                                       463
caccagactt caagaattta nggccatctt tcccgggtac ctg
      <210> 408
      <211> 588
      <212> DNA
```

```
<213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (588)
      \langle 223 \rangle n = A,T,C or G
      <400> 408
                                                                           60
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                                                                          120
agetetetta ttgttateeg aggteaagag getgeaactg teaaggggat gtteteacea
aaagggggtt tgggggaaga ggacacacac aaagctaata aaaccagaat ccccatcccc
                                                                          180
acaaaactca tgggaacaaa atttaaagga taaaacaaaa cccaccaaga cccatattac
                                                                          240
                                                                          300
aaaccaatat ggtaacctgt gttcccttct atggtatgat tatgtcatgt taccttagtg
                                                                          360
ttaaaagatt aacataagga aactgcagca atatataaaa gatatattct ctatagagca
tatttcgatt gattccatta aaataatgac attagaattc catcatangg ttaaaaccag
                                                                          420
                                                                          480
gacaatactg nttttncttt atttaaaaaa aactaccacc taatgactgn attggtcata
acctgaatgg tgtgcaatgg gctcttccat gaatggctgg cngaaacaag cttgggncct gcttgagttt cagctttcct ctttaattta gtngctcaat gataaaca
                                                                          540
                                                                          588
      <210> 409
      <211> 612
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (612)
      \langle 223 \rangle n = A,T,C or G
      <400> 409
                                                                           60
ggtacaaaga tetgacatgt cacccaggga cocatttcac ccactgetet gtttggccgc
                                                                           120
cagtettttg tetetetet cageaatggt gaggeggata ceettteete ggggaagaga
aatccatggt ttgttgccct tgccaataac aaaaatgttg gaaagtcgag tggcaaagct
                                                                           180
gttgccattg gcatctttca cgtgaaccac gtcaaaagat ccagggtgcc tctctctgtt
                                                                           240
                                                                           300
ggtgatcaca ccaatttttc taggttagca cctncagtca ccatacacag ggtaccagtg
tcnaacttga tgaaaatcaa gtaatcnigg ccagtcicta aaatcaaatc tigaatggta
                                                                           360
tcaattcacc cttgatgaag gggaatcggg ggtaacccgg atgggtgccg ggccttnatg
                                                                           420
aagtcancca natgaaggga ttcctttggg gcccccaaag aacttttttn attttcacaa
                                                                           480
                                                                           540
cttgnacctt geeggeggg eegtteaaaa gggenaatte cagneacttg gnggeegtet
                                                                           600
aanggatcca acteggacca acttggegna anatggeaaa etggtteetg gggaaatggt
                                                                           612
atccctccaa tn
       <210> 410
       <211> 353
       <212> DNA
       <213> Homo sapiens
       <400> 410
                                                                            60
acgcggaagc agtggtaaca acgcagagta acgcgggatg gcacatgcag cacaagtagg
                                                                           120
totacaagac gotacttocc ctatcataga agagottato acctttcatg atcacgooot
cataatcatt ttccttatct gcttcctagt cctgtatgcc cttttcctaa cactcacaac
                                                                           180
                                                                           240
aaaactaact aatactaaca totoagacgo toaggaaata gaaacogtot gaactatoot
```

geoegecate atectagtee teategeest eccateceta egeateettt acataacaga

300

```
cgaggtcaac gatccctccc ttaccatcaa atcaattggc caccaatggt acc
                                                                           353
      <210> 411
      <211> 612
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (612)
      <223> n = A, T, C \text{ or } G
      <400> 411
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                                                                            60
                                                                           120
cgtcggctgc tgggaagatc tggattctcg tttcaggtca ccatcagaaa agctaagttt
                                                                           180
qctqtataqt qaqqatcaqq agatctgatc ctgattgcag aaccttccct gattacagaa
tettgggttg tateteccae tteaccette tagaceatee cagaagatet ataagattte
                                                                           240
atctgggaaa tcactaggag ttcttggaag ggaaagaagg aagattgttg gttggaataa aaacagggtt gaatgagttc cagaaagcnn ggttctcaac ctcgtggaca gcaatctgca
                                                                           300
                                                                           360
gaagangaga acttcaaaaa accnactana agcancttgc anagaagtaa aatgagaagg
                                                                           420
ggncttctna ngaaagaaga cacttggncc acagcagaaa aaactttgac cnantnttnc
                                                                           480
caggaagana gggggggtcc cncttttaaa naaccccctt taagatncng gnggaanacc
                                                                           540
tcanngacca ncentaaatt nnggaaaccg aaaaggggen gteetttttg ntnneagntg
                                                                           600
                                                                           612
cnccnttaan nt
      <210> 412
      <211> 607
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (607)
      <223> n = A, T, C or G
      <400> 412
                                                                            60
acgegggget ctctcgccag gegtectegt ggaagtgaca tegtetttaa accetgegtg
                                                                           120
gcaatccctg acgcaccgcc gtgatgccca gggaagacag ggcgacctgg aagtccaact
actteettaa gateateeaa etattggatg attateegaa atgitteatt gigggageag
                                                                           180
acaatgtggg ctccaagcag atgcagcaga tccgcatgtc ccttcgcggg aaggctgtgg
                                                                           240
                                                                           300
tgctgatggg caagaacacc atgatgcgca aggccatccg agggcacctg gaaaacaacc
cagctctgga gaaactgctg cctcatatcc gggggaatgt gggctttgtg ttcaccaagg
                                                                           360
aggacctcac tgagatcagg gacatgttgc tggccaatna ggtgcccagc tgctgcccgt
                                                                           420
                                                                           480
gctggtgccc atttgcccat gtgaangtca cttgtgccca gcccaaaaca cttgtcttng
                                                                           540
ggcccganaa gaacttcttt tttccaggcn ttaaaatatt caccccttaa antttcaagg
                                                                           600
ggccccattt gaaatcctgg annatnngca ttgatcaana ttganacaaa gtggnancnt
                                                                           607
ccaaccc
      <210> 413
       <211> 606
       <212> DNA
```

<213> Homo sapiens

```
<220>
      <221> misc_feature
      <222> (1)...(606)
      <223> n = A,T,C or G
      <400> 413
                                                                           60
acaggicaga gicticitti citticitti tgagatggag tcttgctctg tigccagact
ggagtgcagt ggtgcgatct gggctcactg caatctccac ctcccgggtt caagcgattc
                                                                          120
                                                                          180
tectgeetea geetecegag taaetgggae taeaggtgtg egecaecaag eccageteat
ttttgtattt ttagtagaga tggggtttca cgatgttggc taggatggtc tcgatctctg
                                                                          240
gtcagagtct tttctgtaaa tatccttggt aaagaagcaa ttttagactg tagctgttgc
                                                                          300
aaatgcttta aggaagaagc anaacaactg tcagtcttcc tgaaatgaag aaactacacc
                                                                          360
                                                                          420
agggetgeta tateagagea acceeaacea geacteeaat catgatgeee gacagtggee
                                                                          480
ccagettgag aaccagagaa gttccagatg cagagactgt gagetentga ctatgggaat
tttngnggcn ntaacccaan tttgagacna aacnaggcct tngncccggt tttnatttgg
                                                                          540
                                                                          600
gngggatttt gcggataaan aaacttgnng gggntnctgc ggnatccatg gaacnccaaa
                                                                          606
anatng
      <210> 414
      <211> 624
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (624)
      <223> n = A,T,C or G
      <400> 414
ggtactittt tittitttt tittittttg tagatgaggt cicgctatgt tgcccaggct
                                                                           60
                                                                           120
ggagtgcagt tattcacagg tgcaaccaca gggcactgca gctttaaact cctgggctca
                                                                           180
agegatecte etgeeteage etcecaaata gttgggacta gatgeaegea enaceaegee
tgactcagga cattattett aaaggtatta teeaggaaac agataaggte atteataaaa
                                                                           240
                                                                           300
cacacggntt ttttctttag ctcagtgtta acaatgaaag tagattccac tattgaagca
                                                                          360
caagttgcaa attggtaaca tagngaacat attgntgtag gaaagggggt tcagtgtgnt
gtgttatatn agenettgaa etttttatgg gngtnataag cenngttate ttgneecaaa gaaanneeat tttnaggatt ngatggtttt ettannggaa nannetnggg ggnattntgt
                                                                           420
                                                                           480
ngggcatgaa cttttatgtn ggaatcagtc ccatanaggt aaggggtttn aatcccaaaa
                                                                           540
                                                                           600
ancggggnet tttatgggaa atnnecttta etteaaagge caaanngatn gtnggtgtea
                                                                           624
cttcnaantt ccnqannnca annq
       <210> 415
       <211> 609
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(609)
       <223> n = A,T,C or G
       <400> 415
```

acgogggtta caacggaagt aaaatctgtc gaaatgcacc atgaagcttt gagtgaagct

60

```
cttcctgggg acaatgtggg cttcaatgtc aagaatgtgt ctgtcaaqqa tqttcgtcgt
                                                                            120
                                                                            180
ggcaacngtt gctggtgaca gcaaaaatga cccaccaatg gaagcagctg gcttcactgc
teaggtgatt atcetgaace atceaggeea aataagegee ggetatgeee etgtattgga
                                                                            240
ttgccacacg gctcacattg catgcaagtt tgctgagctg aaggaaaaga ttgatcgccg ntctggtaaa aagctggaag aaggccctaa attcttgaag tctggtgatg ctgccattgt
                                                                            300
                                                                            360
tgatatgggt cotggcaage coatgtgttg ttgagagett teteagacta tecacetttg
                                                                            420
ggtngctttg ctggtcgtga natgagacag acaggtgccn gtggggtggc atcaanncat
                                                                            480
gggacaanaa aggcttnttg gancttgcaa aggtncncaa nttttgncca naagcntcaa
                                                                            540
aagntaattg aatttttccc ctannncctg cnccencttt tannanggnn ggaaaacggc
                                                                            600
ttaaanntt
                                                                            609
      <210> 416
      <211> 577
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(577)
      <223> n = A,T,C or G
      <400> 416
ggtacgagct gattgggaac gggctccaat ggacatggct ctgcagtcaa aatagttagc
                                                                             60
agatggacag gtttggaaaa tgtgagggcc catatcatca tanccagcaa taaggagacc
                                                                            120
aacaccatat ggtctccggc catatccgtt gtgttggtat ctgggtcttg cttccaatta
                                                                            180
gagatacaag actgagacac aggcagtggt ctatcgaata caaatctgga atncaaacac
                                                                            240
tectgaegea taaaattaca taacagneta geatnaneag taageeeeeg caattgagat
                                                                            300
accaatatgg ttgtcaacat ggagaatttt tttctgatga cctgccaact cttgatttgc
                                                                            360
gcccttttca atgcnaaccc aaaactggca tgaagntttt gnatttcaga ccancctgnt
                                                                            420
ggctgnacct tggcttaaca ggtttccatt ggcntatttc natttggatn aantcttgcc
                                                                            480
cntggggggn ttcnaancta ggggccatca nttggtcaaa ctqntttnta aaccatgggq
                                                                            540
genggeteng geettggttg etggenteaa caaaaan
                                                                            577
      <210> 417
      <211> 570
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(570)
      \langle 223 \rangle n = A,T,C or G
      <400> 417
ggtactaaga atattagaga actggaaatc cagttttttt gtggtttttt aagaaagaga
                                                                             60
atctgactcc attgcccagc ttggagagca gtggtgcaat agctggggct acaggcgtga
                                                                            120
gccaccacac caggcctgga aacccagttt taatttgtga actacaaatg gttggcaact
                                                                            180
gatteettaa ttgttattge aggagtagge ecaacatgag tecatatgta gteettetet ggtetggtgg gaactgtggg aaatggtgat gaeegtgaet tgaaataetn agaaggtgea
                                                                            240
                                                                            300
tgacaaacaa attccaagta ttccatcttc cttggaagat cttcctctgg ccctatgata
                                                                            360
taggaageng gaateaaatt tgggetettg ggetaagant aggggtatgg aatgageeee
                                                                            420
egtnaantgg cttgnactte ttettegeta atactgggee etggattaaa acettttgat
                                                                            480
ttnancnata gntagggctt tccttcttgg ttaatcaatt cccagaaacc aacattccca
                                                                            540
```

```
570
atttgggtaa natactccct tgtanaaaaa
      <210> 418
      <211> 570
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(570)
      \langle 223 \rangle n = A,T,C or G
      <400> 418
ggtacttcta cacatctgcc taacttggga atgaatgtgg gagaaaatcg ctgctgctga
                                                                         60
gatggactcc agaagaagaa actgtttctc caggcgactt tgaacccatt ttttggcagt
                                                                        120
gttcatatta ttaaactagt caaaaatgct aaaataattt gggagaaaat atttttaag
                                                                        180
tagtgttata gtttcatgtt tatcttttat tatgttttgt gaagttgtgt cttttcacta
                                                                        240
attacctata ctatgccaat atttccttat atctatccat aacatttata ctacatttgt
                                                                        300
aagagaatat gcacgtgaaa cttaacactt tataaggtaa aaatgaggtt tccaagattt
                                                                        360
aataatetga theagttett ghtatttee aatagaatgg gaetnngnne tgttaangge
                                                                        420
ttaagganaa agggaagata agggttaaaa gttggttaat ggacccaacc ntttnaaaga
                                                                        480
aatgenntan anaatanttt natgantaaa naaaggtnee ingeeengge eggeegtttt
                                                                        540
aaangggcca atttcnagca cnctnggcgg
                                                                        570
      <210> 419
      <211> 574
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(574)
      <223> n = A.T.C or G
      <400> 419
ggtacacctt tgactacagc tgcagaagtg ttcctttaga caaagttgtg acccatttta
                                                                         60
ctctggataa gggcagaaac ggttcacatt ccattatttg taaagttacc tgctgttagc
                                                                        120
tttcattatt tttgctacac tcattttatt tgnatttaaa tgttttangc aacctaagaa
                                                                        180
caaatgtaaa agtaaagatg caggaaaaat gaattgcttg gtattcatta cttcatgtat
                                                                        240
atcaagcaca gcagtaaaac aaaaacccat gtatttnact tttttttagg attttttgct
                                                                        300
ttetgtgatt tttettnttt tttgataett geetaacatg catgtgetgt anaantnagt
                                                                        360
taaccaggga aataaccttg ngatnatggc ctanctttta gtttangtct tatgaanttt
                                                                        420
tcattgacca attctaanca ataatggttt annaacaccg tgntntnaaa atttctggta
                                                                        480
anttggaaat aaaaggtttn nttgaaatgg gccttttcca cnnactttnt ttnncagctn
                                                                        540
tttcttggna aataagccct nggttcctga aacc
                                                                        574
      <210> 420
      <211> 573
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
```

```
<222> (1)...(573)
       <223> n = A,T,C or G
      <400> 420
                                                                                 60
acctccggta gaattcggtg aatccatctg gtcctggact ctttttggtt ggtaaactat
tgattattgc cacaatttca gctcctgtta ttggtctatt cagagattca acttcttcct
                                                                                120
ggtttagtct tgggagagtg tatgtgtcga ggaatttatc catttcttct agattttcta gtttatttgc gtagaggtgt ttgtagtatt ctctgatggt agtttgtatt tctgtgggat
                                                                                180
                                                                                240
eggtqqtqat acceettia teattittta ttgnqtetat ttgattette tetettitt
                                                                                300
taintagict tgctagcagt ctatcaattt ntgtngatcc ttitcanaaa aaccengete
                                                                                360
ctggaattca tttaatnttt tnaaggggtt ttttngtggc ctctaatttc cttcaagttc
                                                                               420
tggetetgat ttaagttaat atneetgget ttttggetae nttttgnaan gnggttggen
                                                                                480
cntgnntttt ctanntcctn ttnaantggg gatngnttnn aangcccatt ttnggaannt
                                                                                540
                                                                                573
tcccgctttn ntttgggggg catttangtt nnn
      <210> 421
      <211> 582
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(582)
      \langle 223 \rangle n = A,T,C or G
      <400> 421
ggtacgcggg ggtccgccat ttcgtggacg ccgggtgagt gagagagttg gttggtgttg ggccggagga aagcgggaag actcatcgga gcgtgtggat ttgagccgcc gcattttta
                                                                                 60
                                                                               120
accetagate tegaaatgea tegtgattte tgteeattgg actgtaaggt ttatgtagge
                                                                               180
aatcttggaa acaatggcaa caaqacqqaa ttggaacqqq cttttggcta ctatggacca
                                                                               240
                                                                               300
ctccgaagtg tgtgggttgn tagaaaccca cccngctttg cttttgntga atttgaagat
ccccgagatg canctgatgc aatccgagag ctanattngn angaacacta tgtggcctgc
                                                                               360
ccgtgtnagg aattggaact ggccgnaatg gttgaaanaa agaangttcg aaaattcgtg gncctncntt ccttttggng gtcgtcngnc cttnagaatg attaatcgnn nggaaggang
                                                                               420
                                                                               480
tectteence tennecenan antitucant aaangaanaa agettttitt ngcaacegn
                                                                               540
aancaggtcc ctttttttag attggganaa atagnngagn tc
                                                                               582
      <210> 422
      <211> 570
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (570)
      <223> n = A,T,C or G
      <400> 422
ggtactctga ggctttagat tcagtttggg tctttggggg ggacctctat catcacgcct
                                                                                60
ataatcatcc cgagagtaat catctctgga gctccacgac cgatcatccc gtctgtcata
                                                                               120
teggtettea tageggtece egecteetet gtagteatea teeetgegat acceaetgee
                                                                               180
aaatgetett etgecactge etateeggga ateatageet etateatagt etetgetgee
                                                                               240
teggteatea tagegatece ggecaceata tegatecata teceggegtg ggecatecga
                                                                               300
```

In a management

```
tacccatccc gatacccatc ccgataccgg ctgaatcata acgatetcga tacttgnetc caaagctate atcacetett ctaggtgggt aagtcatcaa agetgtetgg tagcaaggae
                                                                          360
                                                                          420
                                                                          480
gaagcccttc aagtctggat ctggtttggg cagaatnccc atttttatca cnggccaaaa
gnaacgaatc atccctnggc tttaaccnng ngcttgatcn agcaacgtcc acntcgaaat
                                                                          540
                                                                          570
tntcctnqtt acctananaa ctcttcattg
      <210> 423
      <211> 584
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(584)
      \langle 223 \rangle n = A,T,C or G
      <400> 423
accegggtgg ttaaactteg canaatgeet agatattate etaetgaaga tttgeetnga
                                                                           60
                                                                          120
aagctgttga nccacggcaa aaaacccttc agtcagcacg tgagaaaact gcgagccanc
attaccencg ggaccattet gateatecte actggacgee acaggggcan gagggtggtt
                                                                          180
                                                                          240
ttnctqaaqc agctqqctaq tggcttatta cttgtgactg gacctctggt cctnaatcga
gttcctctac naagaacaca ccaataaatt tgtcattgcc acttcaacca anantcngat
                                                                          300
atcagcaatg taaaaatncc aaancatctt actgatgctt actttaagaa gangaagctg
                                                                          360
                                                                          420
engaageeca anacanenng gaaggtgaga tetttegaca canaagtatg agaanttatg
                                                                          480
agatttacqq agcaanqcan qgattgatca nganaagctt ngggcctcac caaatttttn
nccaanannt tcaaagttta ttttcntnag tttcnnnggg cttncttgcn antctggggn
                                                                          540
tggctttgnc ctaatgggaa tttattnctc ccaaaaatgg nggn
                                                                          584
       <210> 424
      <211> 547
      <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(547)
       <223> n = A,T,C or G
       <400> 424
actcttggtt tgtcaatggg actttccagc aatccaccca agagctcttt atccccaaca
                                                                           60
                                                                          120
tcactgtgaa taatagtgga tcctatacgt gccaagccca taactcagac actggcctca
                                                                          180
ataggaccac agtcacgacg atcacagtct atgcagagcc acccaaaccc ttcatcacca
gcaacaactc caaccccgtg gaggatgagg atgctgtagc cttaacctgt gaacctgaga
                                                                          240
ttcagaacac aacctacctg tggtgggtaa ataatcagag cctcccggtc agtcccaggc
                                                                          300
                                                                          360
tgcagctgtc caatgacaac gggaccctca ctctactcag tgtcacaagg aatgatgtag
                                                                          420
gaccetatga gtgtggaate caqaacqaat taagtgttga ccacagegae ccagtcatte
tggaatgncc tctatggncc aaacgaaccc caccatttcc cctnatacac taattaccgn
                                                                          480
                                                                          540
ccaggggtga accttaagct tttctggcat gcagccttta cccacctggc acagtattct
tggctgn
                                                                          547
       <210> 425
       <211> 567
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<212> DNA

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<213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(567)
      <223> n = A, T, C or G
      <400> 425
                                                                              60
ggtaccatcc tttaatagat ctcatacacc agaattcaga tcatgaatga ctgacagaat
                                                                             120
attttgttgg gcagtcctga tttaaaacta agactggctt gtggttaaat gaatatgttc
agtttttgaa ttttaatagt aactccaatt cagtaaatgg tatcactgtt tacccctttt aaagatatga ttagacttcg ttagtaatgt tcaacttttc acaaagatgg tgagtgccat
                                                                             180
                                                                             240
cttaaaactt actggagatt ggctttatat ttagatttat ataactggtt atgtgaatat
                                                                             300
atttaaatac tggggaaatt gcttcactgt cttagaacca agcaagattc acctgtgttt
                                                                             360
tgtgttcatg ttcatttgcc tcttaaaggc aaggggttga agataaataa ggtagcaatg
                                                                             420
                                                                             480
totataqttt tqqccttaac ctatqccaat cctaattata attccctgga nttnaaaang
                                                                             540
gttnctttta ccttatttgg aanggcnttt taaatngngg gttnntgggn naatatttaa
                                                                             567
aggattattc acccctttca catnttn
      <210> 426
      <211> 563
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(563)
      <223> n = A, T, C or G
      <400> 426
ggtacaattt gttcaaggaa tttttgtaga aaaatacgat cctacgatag aagattctta
                                                                               60
tagaaagcaa gttgaagtag atgcacaaca gtgtatgctt gaaatcttgg atactgcagg
                                                                             120
                                                                             180
aacggagcaa tttacagcaa tgagggattt atacatgaaa aatggacaag gatttgcatt
agtttattcc atcacagcac agtccacatt taacgattta caagacctga gagaacagat
                                                                              240
tettegagtt aaagacaetg algatgttee aatgattett gttggtaata agtgtgaett
                                                                              300
ggaagatgaa agagttgtag ggaaggaaca aggtcaaaat ctagcaagac aatggaacaa ctgtgcattc ttagaatctt ctgnaaaatc aaaaataaat ggtaatgaga attttttatg
                                                                             360
                                                                             420
                                                                             480
acctantgcg gcaaattacc ggaaaaactt congngcctg ggaaggctng gcaaaaggcc
ttcatggtca gntgcttaat tatnctaaat gccntgganc ttttgaccag gntctgaana
                                                                             540
                                                                             563
actgttgncc aattcaacag ggg
      <210> 427
      <211> 567
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (567)
      <223> n = A, T, C or G
      <400> 427
ggtacttttt ttttttttt tttttttt tttttgttaa aaaccataca tccttttat
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120
tgntaagtca taaagaggta tcaaaattaa aagcaaaaat tacagggtaa gacttaacaa
aactactagg agcgtcaaag gaagtgaaaa tgggactagg cgcggggcaa tatgaattaa tgaacatggg aaggacaagg atgggganaa cggtgagcat gtgctgaana tactagggga
                                                                               180
                                                                               240
gaggatetgg tgaaaaattt gatettanae aagegeetag gtaaagaaat aatgggataa
                                                                               300
gatttctaaa coccactatg gagettaaga gteateetng ceattggege tgtetetgne atceteteet teeteaagne tettttteat catnetttga teeaatteea getgggeaat
                                                                               360
                                                                               420
                                                                               480
tececegate tttnattate atcateatte cantanggnn ccenttetta ggaanngntn
ttttggnccc cccttaanat ttcaatttcc cttnnnccca tttttttan ggagnttgtg
                                                                               540
                                                                               567
gcnntggccc ttttnggntt aaaaatn
       <210> 428
       <211> 578
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (578)
       \langle 223 \rangle n = A,T,C or G
       <400> 428
                                                                                60
ggtaccctat gaacctgact ctgtggtcat ggcagaagct cctcctgggg tagagacaga
tettattgat gttggatnea cagatgatgt gaagaaagga ggeeetggaa gaggagggag tggtggette acageaceag ttggtggace tgatggaacg gtgeeaatge ceatgeeeat
                                                                               120
                                                                               180
geocatgeet atgecatetg naaatacnge ettteteata teeactgeea aagggaceat
                                                                               240
canatttcaa tggactgcca atggggacct atcaggcctt tnccaatatt catccacctt
                                                                               300
                                                                               360
cagataccag cnactecece atequatqua tetquanatq acattaatge tgataatgaa
                                                                               420
tatctctttn tgcacanatt gttggtcctg gaccccagcc aanaancctt tgcaaanctt
                                                                               480
netttecaga cetggaggat taettatnga cacenttgte ectaaceaga agttgneeat
ttgngcceng aacancactt teccaactgg canttngetg gateceagnn cettenggat
                                                                               540
ttggaanaac nttggctttt gatggatttt ttccccgg
                                                                               578
       <210> 429
       <211> 572
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(572)
       <223> n = A,T,C or G
       <400> 429
ggtaccaaga gtttgctcct ggctgctttg atgtcagtgc tgctactcca cctctgcggc
                                                                                60
                                                                               120
gaatcagaag cagcaagcaa ctttgactgc tgtcttggat acacagaccg tattcttcat
cctaaattta ttgtgggctt cacacggcag ctggccaatg aaggctgtga catcaatgct
                                                                               180
atcatettte acacaaagaa aaagttgtet gtgtgegeaa atecaaaaca gaettgggtg
                                                                               240
                                                                               300
aaatatattg tgcgtctcct cagtaaaaaa gtnaagaaca tgtaaaaact gtggcttttt
                                                                               360
ctggaatgga attggacata gcccangaac agaaagaacc ttgctgggct ggaggtttca
cttgcacatc atggaagggt ttagtgctta atctaatttg ggcctcactg gacttngncc
                                                                               420
atttaatgaa gttnantcat tattgnnatc atagtttgct ttgtttnaan ccttnncatt
                                                                               480
taaagttaaa actqqaattt nanqqtaatt tnaacttqta nqqtttcctq ggtttagctt
                                                                               540
                                                                               572
tttaaatcnt aatttttcca taagcntttt tg
```

```
<210> 430
      <211> 591
      <212> DNA
      <213> Homo sapiens
     <220>
     <221> misc_feature
      <222> (1) ... (591)
      \langle 223 \rangle n = A,T,C or G
      <400> 430
                                                                         60
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                                                                        120
aaqaqaqqct qqqatqaagg gtgcaaagga atagtaaaga aagcatgttt gagatccana
                                                                        180
acaqaataat qqqtaqtaqa qqqaqqtatt gaggatagaa nagtatatgg gtttggcacc
                                                                        240
acqqqqtqqa taggcaaaac atttggttga taangcgcag attctgaact aacttgtaag
gcttgtctgg ttttaggaca ggtaaaatgg nggaatggta aggagaagtt tataggtttt
                                                                        300
atgageceat getgtanean geaagtgata actngetttt aatecetttt enaaageaat
                                                                        360
                                                                        420
gcctgqnqnt atgaagnata tttggcattt gatcngggtt tnaanggntg attagngttn
ctantqaaca atngnaaagg ggntgccatg atcngtnncc caaggatgng attttanggn
                                                                        480
antetentae ttqtqqqqtt naaqqqtqqn gggnttttae nagqnqqqtc cccnaaqqqn
                                                                        540
                                                                        591
gcctnttggn tntangnaat aaanggccng nnaatngana atccnnnttn n
      <210> 431
      <211> 565
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (565)
      <223> n = A, T, C or G
      <400> 431
                                                                         60
accagtgatg ttttgataca agcatataat gtttaatgat caagtcagga taaatggggt
                                                                        120
atccatcacc tcaagcacat ataatcattt ctttgtatta ggcatattca aattccactc
ttttagttat ttttaaatat ccagtaaatt agatcttatt cattctatct agatgtattt
                                                                        180
ttgtacttta tttttctcaa atatttttac ttatgctttt tgtcattatc cacagtgttt
                                                                        240
                                                                        300
ttttttaaag cctgagccac tttgtggttt cagcctcaat ataataatca tccccttact
                                                                        360
cttagactaa ttccttttcc cctqncactt tgcctgtata ctctgtaaaa atgangacct
                                                                        420
tagaaaatca acatttcctg gtgaactttg agagactatt acaagcagtg cccaaaacag
                                                                        480
tangaataag gcaggtaaaa ccagttggga tagccagatn tattattgat ctggtnggac
aaanggataa nttggngggc atggtttcca nggcantcgn gaattcccca ttagctttaa
                                                                        540
gggtenatnn angntggccc anggg
                                                                        565
      <210> 432
      <211> 578
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(578)
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<223> n = A, T, C or G
      <400> 432
acgegggggc caeegtggag ageagagege ggeggetgga agetgetaag teagageege
                                                                           60
                                                                          120
gatgtteegg attgagggee tegeacegaa getggaceeg gaggagatga aaeggaagat
gegegaggat atgateteet ceataeggaa ettteteate taegtggeee teetgegagt
                                                                          180
                                                                          240
cactccattt atcttaaaga aattggacag catatgaaga caggacatca catatgaatg
                                                                          300
caccgatatg aagageetgg ttacagttte gacteetete tgnaagtgaa taggeecaga
aaggtgtaag agactettig aatggacata aaattetget tgttnagaac caagttttgg
                                                                          360
ntctgggtna ctgacctttc aaaagctaaa attttaaaac tattttgggg aagtttttta
                                                                          420
tttnnntatt nntengtttn ttnataaaaa agtacettgg tneeggnace accenttaag
                                                                          480
ggccnaattn cagnennent ngngggeegn ttaetttnng ggatnentaa nttegggane
                                                                          540
                                                                          578
cnaancttgg ggggtaantc angggtcata nnctggtt
      <210> 433
      <211> 563
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(563)
      <223> n = A, T, C \text{ or } G
      <400> 433
acticitictg gccaaagget gttccacatt cactacattt aaaaggette tetecaatat
                                                                            60
                                                                           120
ggattttctc atgctcagta aggttggatt tgccactgaa ggtttttcca cactccttac
atacaaaggg ctictctcct gigtgagttc tctggtgict gatgaggttt gacttctgaa
                                                                           180
tgaaagettt eeegeaatet ttacaeteaa aaggttttte teeagtgtga attttetggt
                                                                           240
gcgtaaggag gttttccttc tggctaaatg attttccaca ttcattacat tcgaaaagct tctcgccagt atgggtgttc tgatgtttaa tgacatactg cttttggcta aaggcttttc
                                                                           300
                                                                           360
cacactegtt acatteaaaa gggttetete teegtgtgaa aatgeteatg eteantgang
                                                                           420
tttgaattgn nggcttgaag acttttccca tacccttaca ggcaaanggg gttttccccn
                                                                           480
ttggaanatn tntggctgcn tnaagntggt gacatctgga inggaaacct tttccncatt
                                                                           540
tccaaaggnn tttttttcnn nag
                                                                           563
      <210> 434
      <211> 563
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(563)
      <223> n = A,T,C \text{ or } G
      <400> 434
ggtacagctg tetgeattga aaatteatge atggagaaag ggagtaagea agggagaaae
                                                                            60
ggtgcgattc acatattccg cgagatcatc aagccagcag agaaatccct ccatgaaaag
                                                                           120
ttaaaacaag ataagegett tageacette eteageetae ttgaagetge agaettgaaa
                                                                           180
gageteetga cacaacetgg agactggaca tratttgtge caaccaatga tgettttaag
                                                                           240
ggaatgacta gtgaagaaaa aqaaattctg atcqqqacaa aaatqctctt caaaacatca
                                                                           300
ttctttatca cctgacacca ggagttttca ttggaaaagg atttgaacct ggtgttacta
                                                                           360
```

```
acattttaaa gaccacacaa ggaaacaaaa tettttettg aaagaaagta aatngateea
                                                                        420
cttctggtga atgaatttga aattcaaagg aatctggcct tcatgccanc aaatgggggt
                                                                        480
                                                                        540
aattcatgnt ggagaataac ctcctttatc cagccgnaca cacctgttgg aaatggatcc
                                                                        563
aactgctgga aattncttaa taa
      <210> 435
      <211> 558
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(558)
      <223> n = A, T, C or G
      <400> 435
ggtacgcggg ggaagatggc ggccgtgcag gcggccgagg tgaaagtgga tggcagcgag
                                                                         60
                                                                        120
ccgaaactga gcaagaatga gctgaagaga cgcctgaaag ctgagaagaa agtagcagag
aaggaggcca aacagaaaga gctcagtgag aaacagctaa gccaagccac tgctgctgcc
                                                                        180
                                                                        240
accaaccaca ccactgataa tggtgtgggt cctgaggaag agagcgtgga cccaaatcaa
tactacaaaa tccgcaqtca agcaattcat cagctgaagg tcaatgggga agacccatac
                                                                        300
ccacacaagt tccatgtaga catctcactc actgacttca tccaaaaata taagtcacct
                                                                        360
gcagcctggg gatcacctga ctgacatcac cttaaaggtg gcaggtagga tccttccaaa
                                                                        420
                                                                        480
aganctintg ggggaaacin antititint tgaactitica aggaaanggg tgaagtitige
agtcatgggc caattccaga aattttaaat cagnagaaga atttttccta ttaataccaa
                                                                        540
                                                                        558
ctgggtcggg ggagactn
      <210> 436
      <211> 528
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (528)
      <223> n = A,T,C \text{ or } G
      <400> 436
ggtacaaaaa aaaccttaca taaattaaga atgaatacat ttacaggcgt aaatgcaaac
                                                                         60
                                                                        120
cgcttccaac tcaaagcaag taacagccca cgatgttctg gccaaagaca tcagctaaga
aaggaaactg ggtcctacgg cttggacttt ccaaccctga cagacccgca agacaaaaca
                                                                        180
actggttett gccagcetet agagaaatee cagaacacte agecetgaca egttaatace
                                                                        240
                                                                        300
aaggggaaca gttaactcca atacaaggtc aaaatcagca acaagttcta caatccagtg
ctgatatcag atacaaagct tcaagggcaa tttcttttcg aaggcttatt ccagtttcgt
                                                                        360
gaggctagca tgaagtgtgt gcatttgcca ggggcaaatt tctattctca attaacccat
                                                                        420
                                                                        480
gcagcaaant gctacgcatc tggctgagtc cggtttanaa nccatttgcc ggnggaccaa
                                                                        528
tggaagggc ccgaattcgt cnnaacttgn cccgggcggg ccgttcaa
      <210> 437
      <211> 576
      <212> DNA
      <213> Homo sapiens
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<220>
      <221> misc_feature
      <222> (1) ... (576)
      <223> n = A, T, C or G
      <400> 437
                                                                           60
acttttttt tttttttt ttttttttt aggtttgagg gggaatgctg ganattgtaa
tgggtatgga gacatgtcat ataagtaatg ctagggtgag tggtaggaag ttttttcata
                                                                          120
ggaggtgtat gggttggtcg tagcggaatc gggggtatgc tgttcgaatt cataagaaca
                                                                          180
gggaggttag aantagggtc ttggtgacaa aatatgttgt gtagagttca gggganagtg
                                                                          240
                                                                          300
cqtcatangt tqttcctagg aanattgtac nggtgagggt tgtttattat aataatgttn
                                                                          360
gggtatccgg ctntgaaana atngggccaa ngggcctgcg gtgtattcga ngttnaaacc
                                                                          420
tgagactagt teggacteee ntttgeaagg neceaaaggg ggttnggttt ggeeettget
annggtgnga naataaatcn tntttattgg cccaagggtt cttaacngen aggagtnaat
                                                                          480
                                                                          540
ccaaagggt nontnggntt ttnnnanaaa nggttgnnaa aaggttaaag ggacconcet
                                                                          576
ttntnnntaa tgntcgnaat gtcaaatnga tngcnn
      <210> 438
      <211> 576
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (576)
      <223> n = A, T, C \text{ or } G
      <400> 438
                                                                           60
ggtaccccaa ttaccagtat ggtggaccct accccttctt ctctgcattg ggaaacagaa
                                                                          120
cagagaacag aaaaaatcat tocatottgo tottaactot ttocacotat gtgctcagtt
tttcaagtag aatttctatt cctttgctgg tgcttttggt tttttccaat gtaggaatca
                                                                          180
agetttteag tgeagetttg aetttgtttg caaetteeag gteacaaete tggaggagge tagaaagaat aatggeaeet egatttaeae tageeeagga etteaggtte tteataeeaa
                                                                          240
                                                                          300
catgetetae aagigttitt geaaaacaae ettetetee attnietti catettita
                                                                          360
tettgeteta ttaaccaett nagaaactaa gaatgteeet geaaggatgt tetggeaatg
                                                                          420
                                                                          480
ntgaaagett eteegteett ggeeaceagg atgeaagtee ntggtinttg ceagettgge
                                                                          540
cnatnggcat tecatngqna nggettgaac cgttttccag ggggcagant cccaaaatgg
                                                                          576
cengacacca accenacang cagacttntt ttagen
      <210> 439
      <211> 578
      <212> DNA
      <213> Homo sapiens
      <220>
       <221> misc feature
      <222> (1)...(578)
      <223> n = A,T,C or G
       <400> 439
cgaggtacgc gggggagaaa aaacctgcgg aaaatggtag cgatggcggc tgggccgagt
                                                                           60
                                                                          120
gggtgtctgg tgccggcgtt tgggctacgg ttgttgttgg cgactgtgct tcaagcggtg
                                                                          180
tetgettttg gggeagagtt tteateggag geatgeagag agttaggett ttetageaac
```

```
240
ttgctttgca gctcttgtga tcttctcgga cagttcaacc tgcttcagct ggatcctgat
tgcagaggat gctgtcagga ggaagcacaa tttgaaacca aaaagctgta tgcaggagct
                                                                               300
                                                                               360
attettgaag tttgnggatg aaaattggga aggtteeetn aagteeaane ttttgttang
agtgataaaa cccaaactgt tcagaaggac tgccaaatna aagtatgtnn cgtggtttca
                                                                               420
                                                                               480
aaccntgaat taaaaggctt ttngaccaac atngggnaca attgcttgan nacttgtcca
tttcttaaaa ttgggaacnc tggaccnggt nanaaanatt tcngattgga aaantttgga
                                                                               540
                                                                               578
concatttta aatottgott aaattttggo caatcott
      <210> 440
      <211> 573
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
       <222> (1)...(573)
      \langle 223 \rangle n = A,T,C or G
      <400> 440
ggtacttttt ttttttttg agacagggtc ttgccctgtc acccaggctc gagtgcactg
                                                                                 60
gagtgatcac agetcactgg cetcaagtga teeteetgee ttggeceett aagtgecagg
                                                                               120
gttacaggca tgagctacca tgcctggcag aaattcaaga tttggataaa cttacttctt
                                                                               180
tgccaageet gttetteaag ttatteagaa etgggtgtat acettgteet catatgtate
                                                                               240
                                                                               300
ttgtccctgc tgtcttttag gttagcaagg tgtatgaata cttttaagtt ttgtttgttc
                                                                               360
ttttcctcgt ggtatcaagt gaaatactga tctattctct ggctagggtc aatttacaaa
                                                                               420
attgccatgg aactgagcca aaaggcccca cgtgggataa aaattnctta ccatcgacgc
                                                                               480
ccancegtan tttttcaagg tattggettt tggaagnttt accaaattte nggtaaacca
                                                                                540
aaattchaaa agnaaaaaat theeetggng taacettgee egggeggeeg tteaaaaggg
                                                                               573
cnaatttcca ncacattggg cggccgttaa tna
       <210> 441
       <211> 572
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1)...(572)
       <223> n = A, T, C \text{ or } G
       <400> 441
                                                                                 60
ggtacaaaat tttattaaag gtctttagag agcaacatcc agactccaga atacagctgc
caaggagace etgttatget gtggggactg getggggeat ggeaggegge tetggettee
                                                                                120
caccettetg ttetgagatg ggggtggtgg geagtatete atetttgggt tecacaatge teaegtggte aggeaggge ttettagge caatettace agttgggtee cagggeagea tgatetteae ettgatgee ageacacet gtetgageaa caegtggege acageagtgt
                                                                                180
                                                                                240
                                                                                300
caacgtagta gttaacaggg gtctccgctt gtggatcatc aagccatcca caaacttcat ggatttagcc ctctgncctt cggaggttcc cagacaccca caanctngca agcctttggc
                                                                                360
                                                                                420
                                                                                480
cccacttttc catgatgaaa ctgnagncac aaccatangc aagggccctt cggacannta
                                                                                540
aggeetteet aaggagnttg naacnenana naacttttge ttgggeantg ggeacaceag
                                                                                572
nacctntaag nggccccctt tttaagcata aa
```

```
<211> 562
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(562)
      \langle 223 \rangle n = A,T,C or G
      <400> 442
acaggicaga gicticitti citticitti tgagatggag tottgcictg tigccagact
                                                                               60
                                                                              120
ggaqtqcaqt ggtgcgatct gggctcactg caatctccac ctcccgggtt caagcgattc
                                                                              180
tectgeetea geeteeegag taaetgggae taeaggtgeg egeeaeeaag eecageteat
ttttgtattt ttagtagaga tggggtttca cgatgttggc taggatggtc tcgatctctg
                                                                              240
                                                                              300
gtcagagtct tttctgtaaa tatccttggt aaagaagcaa ttttagactg tagctgttgc
aaatgcttta aggaagaagc aaaacaactg tcaagtcttc ctgaaatgaa gaaactncac
                                                                              360
                                                                              420
cagggetget atateagaac aaceneaace aageacttea aacatgatge egacaggtgg
ccccagetta aaaaaccagg aanaagtten ganteeenaa actgngaatg cetettggae ttttggaatt aattggggge cagtageeaa gttatnagae caaatcangg entagggeee
                                                                              480
                                                                              540
cgtattattt ggcggggatt tg
                                                                              562
      <210> 443
      <211> 585
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(585)
       <223> n = A,T,C or G
       <400> 443
                                                                               60
actitiatit titiggiggig aaattgactg atgattitice titticitieg ciggactatt
gtgccaactg ccaggetgcc tectgeeett acageeetaa gtggetgeet tetttecate
                                                                              120
aactcccaac ttcttcctgt gaagtttaat tgtctcaacg cctccccctc ccccattccc tccatttttc tcccaagaaa cctgactcaa ttatttgcat attttgagaa actgctgcag
                                                                              180
                                                                              240
attagttett tttgecagtt tteeetggaa eteetggeet tttgtggagg ggagggatgg
                                                                              300
agagaatagg aatcttcact agaagccgtg ggaagaattg gaagttacat gctgtatatg
                                                                              360
                                                                              420
caatgtccag cagtctgata aactgacgat tcttaatcaa gattttttcc tgatggggaa
                                                                              480
gggactttta ttttctttta nagaggggaa agtgtgagct cttcccttat tcctaatggc
                                                                              540
tatttttgaa gcaaanaagg ccacaacatt ngcacatgcc acctgcnaag gaccttgagt
                                                                              585
nagtgaagnc tcctaaaact gggttaanaa ccttgttttc tctnn
       <210> 444
       <211> 437
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (437)
       <223> n = A, T, C or G
```

```
<400> 444
                                                                               60
acgcggggac gtgactcagc actttcccca gagcccggac tgcggagaac aatatcctcc
tecetaacag ataaacagee ettgtteete gggataagga etggeagtee eetgacaeee
                                                                              120
taagaccggc atctgtcgat gttatttccc cagcatggcc gaaacagaag ccctgtcgaa gcttcgggaa gacttcagga tgcagaataa atccgtcttt attttgggcg ccagcggaga
                                                                              180
                                                                              240
                                                                              300
aaccggcaga gtgctcttaa aggaaatcct ggagcagggc ctgttttcca aagtcacgct
cattggccgg aggaagctca ccttcgacga ggaagcttat aaaaatgtga atcaagaagt
                                                                              360
                                                                              420
ggtggacttt gaaaagttgg atgactacgc ctctgccttt caaggtcatg atgttggatt
                                                                              437
ctgtgcctgg gtacctn
      <210> 445
      <211> 592
      <212> DNA
      <213> Homo sapiens
      <220>
       <221> misc_feature
       <222> (1)...(592)
       <223> n = A,T,C or G
       <400> 445
actititit tittitit tittititt taaggittga gggggaatgc tggagattgt
                                                                               60
aatgggtatg gagacatatc atataagtaa tgctagggtg agtggtagga agttttttca
                                                                               120
taggaggtgt atganttggn cgtagcggaa tcggggggtat gctgttcgaa ttcataagaa cagggaggtt aaaagtaggg tcttggtgac aaaatatgtt gtgtanagtt caggggaaag
                                                                              180
                                                                              240
tgcgtcatat gttgttccta ggaanattgt antggtgagg gtgttaatta taataatgtt
                                                                               300
tgtgtattcg gctatnaana atagggccaa atgggcctgc ngcctattcn atgtttaanc
                                                                               360
                                                                              420
tgagacttnt tcggactccc cttcggcaan gtcnaantgg ggttcggttg ngcnctgcag
                                                                               480
tgnggagata nntcntntta ntggccaatg gtnnngatgg ccagaataat cannanggnt
                                                                              540
tentinnten inaaaaggie naaatggiin angganacen ettattagga attgitaate
ttnaatgatn gttntggnga enetatatgg anaatgtnag gnetaeteen ng
                                                                              592
       <210> 446
       <211> 599
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (599)
       <223> n = A,T,C or G
       <400> 446
ggtacggcaa acacaacgga cctgagcact ggcataagga cttccccatt gccaagggag
                                                                                60
                                                                               120
agegecagte ceetgttgac ategacacte atacagecaa gtatgaceet teeetgaage
                                                                               180
ccctgtctgt ttcctatgat caagcaactt ccctgaggat cctcaacaat ggtcatgctt
tcaacgtgga gtttgatgac tctcaggaca aagcagtgct caagggagga cccctggatg
                                                                               240
gcacttacag attgattcag tttcactttc actggggttc acttgatgga caaggttcat
                                                                               300
agcatactgt ggataaaaag aaatatgctg cagaacttca cttggttcac tggaacacca aatatgggga ttttgggaaa gctgtgcagc aacctgatgg actggccgtt ctaggtattt
                                                                               360
                                                                               420
tttttgaagg ttggcagcgc taaaccnggc cttnataaag ttgttgaatg tgctggattc
                                                                               480
cattaaaaca aagggcaaga attgctgact ttcactaatt nnaatcctcg tnggccttct
                                                                               540
tectgaaate ettggattae eggacetnee eagettaetn accaneette tetttingg
                                                                               599
```

```
<210> 447
      <211> 588
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (588)
      <223> n = A, T, C \text{ or } G
      <400> 447
ggtacgcggg atgagtgtgg aatccagaac aaattaagtg ttgaccacag cgacccagtc
                                                                            60
                                                                           120
atcotgaatg toototatgg cocagacgac cocaccattt coccctcata cacctattac
cgtccagggg tgaacctcag cctctcctgc catgcagcct ctaacccacc tgcacagtat
                                                                           180
tettggetga ttgatgggaa catecageaa cacacaaag agetetttat etceaacate
                                                                           240
                                                                           300
actgagaaga acageggact ctatacetge caggecaata acteagecag tggecacage
aggactacag tcaagacaat cacagtetet geggagetge caagecetee atetecagea
                                                                           360
                                                                           420
acaactecaa accegtggag gacaaggatg ctgtggcctt ccctgtgaac ctgaggctca
gaacacaacc tacctgtggt gggtaaatgg tcagagcctc cagcagtccc aaggctggag
                                                                           480
ctgtccaatg gcaacangga cctnactcta ttcaatgtca caagaaatga cncaagaacc
                                                                           540
tatgnatgtg gaatccagaa ctnagtgatg caaaccgaat gaccagnn
                                                                           588
      <210> 448
      <211> 593
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (593)
      <223> n = A, T, C or G
      <400> 448
accatttgtc tgacctctgt aaaaaatgtg atcctacaga agtggagctg gataatcaga
                                                                            60
                                                                           120
tagttactgc tacccagagc aatatetgtg atgaagacag tgctacagag acctgctaca
                                                                           180
cttatgacag aaacaagtgc tacacagctg tggtcccact cgtatatggt ggtgagacca
                                                                           240
aaatggtgga aacagcctta accccagatg cotgetatec tgactaattt aagtcattge
tgactgcata getetttte ttgagagget etceattttg atteanaaag ttageatatt tattaceaat gaatttgaaa eeagggettt ttttttttt ttgggtgatg taaaacneaa
                                                                           300
                                                                           360
ctncctgnca ncaaaataat taaaatagnc acattgntat cttttattag gtaattcact
                                                                           420
tottaattan atggntcaat actotaagna toaaaatntt coaattatna tggctcacot
                                                                           480
                                                                           540
gaaagaagna tgctctttta aggaatacag cttcttcnat tnacaattta acanggggag
aaaattaaan tnaangantt ganatetgga ggngtannaa ngntetegen tte
                                                                           593
       <210> 449
       <211> 577
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(577)
```

```
<223> n = A, T, C or G
      <400> 449
actgtgggtc gaagtaatgg atacggacgt aaccatcttc gccgccgctg ctgtagctct
                                                                           60
tgccatcagg atggaaggca acactgttga taggtccaaa gtgacccttg actcttccaa
                                                                          120
actettette aaaggeeaaa tggaagaace tggeeteaaa ettgeeaate etggtggagg
                                                                          180
                                                                          240
ttgtggttac atccatggct tcctgaccac cgcccaggac cacatggtca tagttggggg
                                                                          300
agagggcagc tgagttgaca ggacgttctg tccggaaagt cttctgatgt tcaagagttg
                                                                          360
tggagtcaaa aagcttggct gtgttgtcct tggacncggc acaaacatgg tcatgtccct
                                                                          420
ggataactgg atgtcgttga tctgccggga gtgctcctta acattcacca acacctcttc
anacttggca ctatactggt tgactctcca ctcttatggc enggatgatg cactccccca
                                                                          480
aggggtncca aacagnactg gtgatttaga atcattgcan ggatcttatg tagggctcat
                                                                          540
                                                                          577
tgntgcaatc tggcttggat ccgcagtcaa aaaagnt
      <210> 450
      <211> 575
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(575)
      <223> n = A, T, C \text{ or } G
      <400> 450
                                                                           60
ggtacttgtg atcacactac gggaatctct gtggtatata cctggggcca ttctaggctc
                                                                          120
tttcaagtga cttttggaaa tcaacctttt ttatttgggg gggaggatgg ggaaaagagc
tgagagttta tgctgaaatg gatttataga atatttgtaa atctattttt agtgtttgtt cgttttttta actgttcatt cctttgtgca gagtgtatat ctctgcctgg gcaagagtgt
                                                                          180
                                                                          240
ggaggtgccg aggigtcttc attctctcgc acatitccac agcacctgct aagtitgtat
                                                                          300
                                                                          360
ttaatggttt ttgtttttgt ttttgtttgt ttcttgaaaa tgagagaaga gccggagaga
                                                                          420
tgattittat taattnttnt ttttttttt tactatttat agctttaaaa agggccincc
                                                                          480
ttcccctctt ctttctttgg nctctttcat taaccccttc ccagtttttt ttaacttaaa
ccccgttctc atggcctngg ccttttgaag cgnttcctct tataaaaagc tttgccgaac
                                                                          540
                                                                          575
aantttttt taccgatccc aaatttatga agggg
      <210> 451
       <211> 573
       <212> DNA
       <213> Homo sapiens
       <221> misc feature
       <222> (1) ... (573)
       <223> n = A, T, C or G
       <400> 451
                                                                            60
actaggctaa ctagaaggat ctcatcccca tatgtggtct catttcaagt ctatggatga
                                                                           120
ctaccttcat tgctgtgtgc gagatggttt caccccttga aaatatggtc acttcagcat
                                                                           180
aaaatagtta aatctttata atgatcaatt catcctacct ccttttacat gcagctgaaa
aatgacaggc tagggacata gaatattgtg aactttatac tgttagaatc actgtccatt
                                                                           240
                                                                           300
aaatgatcac tagctaatgg tcactaaatt tacaaattaa ggaaattata tatagaatac
tgcaaaaaca cagtaaaaag actgaagttc gcccatttct gctcaggaag tctcttcact
                                                                           360
```

```
cctaagette atatgttgee ttetggette aaaattetge tattattaet gtttteetee
                                                                              420
                                                                              480
tttqatcttc ctttqqtccc caqtqccaqa cttccaagcc ttttnqttaa aaagccatct
                                                                              540
tttggatgcc atttcnaaca gcttcagtga tgcctctgaa aaaaggatct gccggctaan
                                                                              573
atttctcngg ttcgtgcttt ctaccgganc tcc
      <210> 452
      <211> 595
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(595)
      \langle 223 \rangle n = A,T,C or G
      <400> 452
acaattttat ccctaaaact ctgttgacat caaaatatga cagttgctat atccataaaa
                                                                               60
                                                                              120
tatttacata gcacggcata ttaagcttta gacacttggc aattaaacca cataaaaaga
ggacaagace cecatectae atqtttggaa teaggtgtte aceggteeet atetggegae
                                                                              180
tgtacgcggg tggggtcctt acttgtattc tgttatcagc tgaltttgaa acatataata
                                                                              240
atgattttct tgttcccttc tttaactagc tgcctttaga ttttgataat cacagtctta
                                                                              300
aaatactagg aaagaagtgg atgggaattg taggcataga tttcatatca agggcatttc
                                                                              360
aagacagaat ttttaattcc tgtagtaggc ttgctggagc naaaggaaaa tgtgctggtt
                                                                              420
aaaaatcaac ttatgccatt ttaaaatttg ataaaatttg gagtggcatn ctgctaaggg
                                                                              480
gagacettgg geeggaceee ettangggea aatteengea eactgggggg eggtactang
                                                                              540
gggatccgac ntcggnccan acttggcgna tcatgggctt antgttcctt gnggn
                                                                              595
       <210> 453
       <211> 380
       <212> DNA
       <213> Homo sapiens
       <221> misc_feature
       <222> (1) ... (380)
       <223> n = A, T, C or G
       <400> 453
ggtacgcggg gagccgcctg gataccgcag ctaggaataa tggaatagga ccgcggttct
                                                                               60
                                                                              120
attttgttgg ttttcggaac tgaggccatg attaagaggg acggccgggg gcattcgtat
tgcgccgcta gaggtgaaat tcttggaccg gcgcaagacg gaccagagcg aaagcatttg ccaagaatgt tttcattaat caagaacgaa agtcggaggt tcgaagacga tcagataccg
                                                                              180
                                                                              240
tegtagttee gaccataaac gatgeegace ggegatgegg eggegttatt eccatgacee geegggeage tteegggaaa ecaaagtett tgggtteegg ggggagtatg gttgeaaaaa
                                                                              300
                                                                              360
aaaaaannaa aaaaaaaaqt
                                                                              380
       <210> 454
       <211> 589
       <212> DNA
       <213> Homo sapiens
       <221> misc feature
```

```
<222> (1)...(589)
      <223> n = A,T,C or G
      <400> 454
ggtactcttg gtttatcaat gggacgttcc agcaatccac acaagagctc tttatcccca
                                                                              60
acatcactgt gaataatagc ggatcctata tgtgccaagc ccataactca gccactggcc
                                                                             120
                                                                             180
tcaataggac cacagtcacg atgatcacag tctctggaag tgctcctgtc ctctcagctg
tggccaccgt cggcatcacg attggagtgc tggccagggt ggctctgata tagcagccct
                                                                             240
ggtgtatttt cgatatttca ggaagactgg cagattggac cagaccctga attettctag ctcctccaat cccattttat cccatggaac cactaaaaac aaggtctgct ctgctcctga
                                                                             300
                                                                             360
agccctatat gctggagatg gacaactcaa tgaaaattta aagggaaaac cctcaggcct
                                                                             420
gangtgtgtg ccactcagag acttcaccta actagagaca gtcaaactgc aaccatgggt
                                                                             480
gagaaattga cgacttcaca ctatggacag cttttnccaa gatgtcaaac aagactcctc
                                                                             540
atcatgataa ggntcttacc cctttaattg nccttgttat gcctgccct
                                                                             589
      <210> 455
      <211> 589
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (589)
      <223> n = A,T,C or G
      <400> 455
ggtacgcgga agagacaggg tttcaccatg ttgcccaggc tggtttcgaa ctcctgacct
                                                                              60
caggigated accedenced geoteceaaa gigetgggat tacaggetig ageocoegeg cocagecate aaaatgetit trattietge atatgitgaa tactittiae aattcaaaaa
                                                                             120
                                                                             180
aatgatctgt tttgaaggca aaattgcaaa tcttgaaatt aagaaggcaa aaatgtaaag
                                                                             240
gagtcaaaac tataaatcaa gtatttggga agtgaagact ggaagctaat ttgcattaaa
                                                                             300
ttcacaaact tttatactct ttctgtatat acattttttt tctttaaaaa acaactatgg
                                                                             360
atcagaatag ccacatttag aacacttttt gttatcaagt caatattttt agatagttag
                                                                             420
aacctggtct taagcctaaa agtgggcttg attctgcagt aaatcnttta caactgcctc
                                                                             480
                                                                             540
gacacacatt aaccttttta aaaatngacc ttcccgaagt cttttggtag catggnacac
                                                                             589
ctgatgctta natgttcang taattaatat ggnccagnag tnttgtnnc
      <210> 456
      <211> 582
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(582)
      \langle 223 \rangle n = A,T,C or G
      <400> 456
acagaatgtt gatacaaagc ttaaaattct tgcatatggt catagaaaat gcatctttgg
                                                                              60
ttttgtgttt ttatcacttg cttccaactt aggettttgg ctcagaagat tattgaataa
                                                                             120
tgatttgtct tagtttctgt ttcagtaagg gaattctgag gccgttgcta tgataccatc
                                                                             180
attaagacat toacatgtot toatataata totottoatt toaaatoota atcactattt
                                                                             240
```

300

catactatta cagggetttg atgetgecag cactgtettt tacataggaa attetagatt

```
tgcacagtaa tagaggaatt agaagtacct aactatacac tttgattcag cctgctaaat
                                                                        360
caggggttca atactagett ggacaaactt tgtaagtaat taattgetae cageettatt
                                                                        420
ggaaacaaat tatcaactag tttcccctgc caarttttga aattcactgn ttcacttaat
                                                                        480
ctattatatt actaataatg gattaataaa gatgaattaa ttattattac ttactagtnt
                                                                        540
aaatgaaaaa cagggactga aatagtctgn atccgngttg ca
                                                                        582
      <210> 457
      <211> 380
      <212> DNA
      <213> Homo sapiens
ggtacttttt ttttttttt tttttggagt ttttagttta ttaatgttct tgcgaaaaat
                                                                         60
ccacagtggc cacagctaac atcattgcag cacctttact cetteggetg tgatccaate
                                                                        120
tccagetcac ttettttgc cageaccaac attggeettt geagteceec tgaetttett
                                                                        180
cattetgtte ttgcgttect ttcgttgctt tettgaggte tttttettet catacaggee
                                                                        240
atgtettgea agtetatgtt tgggtteatt tttetttgea taateeaggg aateataaat
                                                                        300
Catgccaaag ccagttgtct tgccaccacc aaaatgagtt ctgaatccaa atacaaagat
                                                                        360
gacatccggt gtggtcttgt
                                                                        380
      <210> 458
      <211> 382
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(382)
      <223> n = A, T, C or G
      <400> 458
acgcggggag aacagccacc cctctctcgg gcactgctgc catgaatgcc ttcctgctct
                                                                         60
ccgcactgtg cctccttggg gcctgggccg ccttggcagg aggggtcacc gtgcaggatg
                                                                        120
gaaatttctc cttttctctg gagtcagtga agaagctcaa agacctccag gagccccagg
                                                                        180
agcccagggt tgggaaactc aggaactttg cacccatccc tggtgaacct gtggttccca
                                                                        240
tectetgtag caaceegaac titecagaag aacteaagee tetetgeaag gageecaatg
                                                                        300
cccaggagat acttcagagg ctggaggaaa tcgctgagga cccgggcaca tgtgaaatct
                                                                        360
gtgcctacgc tgcctgtacc tn
                                                                        382
      <210> 459
      <211> 592
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(592)
      \langle 223 \rangle n = A,T,C or G
      <400> 459
ggtactgagg aaatattttg taaagtgagc tttggggtata acttagcccc atcattattt
                                                                         60
agagaataga ggaggaagaa agaggaagga ttttaaaggc agacaatgac agaccattca
                                                                        120
ggataggtag ggttttaaag ggagataaac acagtctcat caactaagga gagatttgct
                                                                        180
```

;----

```
240
gcagtaaata ggatgaggga aatagtctgt gggatgcaag caaaggaagc agggtgcctt
agacactgag tggagccaga aagatcatgc ggcctttttc caagtacatg gccaccaagt
                                                                          300
                                                                          360
aagaatggtt ggtgacaaga cagaaggcta aaacaggaag gtaatcttgt gcacctgaca
aatngaaga attaaggatc aaaattgaag caggctntaa gagtttcaag aaattcttaa
                                                                          420
aacccaaaag tgatttggaa gccccaaact ttccggtaat gctncccatg gcatgatggg
                                                                          480
ccaaaacctt gggggttect aagttnnaaa agccetntne caaattttaa tggacccet
                                                                          540
                                                                          592
acattttttc taatcaatcc cccctttcca aaaaaatngg acctcntttt tt
      <210> 460
      <211> 578
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(578)
      \langle 223 \rangle n = A,T,C or G
      <400> 460
acgcgggcac tatcctgaat tatgtgcctg tctagataag cagagaccat gccaaagcta
                                                                           60
taatggaaaa caagtttaca aagagacctg tatttctttc ataaaagact tcttggcaaa
                                                                          120
aaatttgatt atagttattg gaatagcatt tggactggca gttattgaga tactgggttt
                                                                          180
ggtgttttct atggtcctgt attgccagat cgggaacaaa tgaatctgtg gatgcatcaa
                                                                          240
gctatcgtca gtcaaacccc tttaaaatgt tgctttggct ttgtaaattt aaatatgtaa
                                                                          300
                                                                          360
gtgctatata agtcaggaqc agctqtcttt ttaaaatgtc tcggctagct agaccacaga
                                                                          420
tatcttctag acatattgaa cacatttaag atttgaggga tataagggaa aatgatatga
atgtgtattt ttactcaaaa taaaagtaac tgttacgttg cgaaaaaaan nnnnnnnnn
                                                                          480
naaaaaaaag tneettggge egggaceaeg etagggeaaa teeageaeae tggeggeegt
                                                                          540
actagggatc cactnggacc agctggcgna atatggnn
                                                                          578
      <210> 461
      <211> 425
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(425)
      <223> n = A,T,C or G
      <400> 461
acgcggggct ttctggtctc ggccgcagaa gcgagatgac gaagggaacg tcatcgtttg
                                                                           60
                                                                          120
gaaagcgtcg caataagacg cacacgttgt gccgccgctg tggctctaag gcctaccacc
ttcagaagtc gacctgtggc aaatgtggct accctgccaa gcgcaagaga aagtataact
                                                                          180
ggagtgccaa ggctaaaaga cgaaatacca ccggaactgg tcgaatgagg cacctaaaaa
                                                                          240
ttgtataccg cagattcagg catggattcc gtgaaggaac aacacctaaa cccaagaggg cagctgttgc agcatccagt tcatcttaag aatgtcaacg attagtcatg caataaatgt
                                                                          300
                                                                          360
tctggtttta aaaaatnnan nnnaannntn ntnnaaanaa aaaaagtnct nggccgngac
                                                                          420
cacge
                                                                          425
      <210> 462
      <211> 581
      <212> DNA
```

```
<213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (581)
      \langle 223 \rangle n = A.T.C or G
      <400> 462
ggtactattg acccagegat gggggcttcg acatgggctt tagggagtca taagtggagt
                                                                          60
cogtaaagag gtatetttac tataaaaget attgtgtaag etagteatat taagttgttg
                                                                         120
gctcaggagt ttgatagttc ttgggcagtg agagtgagta gtagaatgtt tagtgagcct
                                                                         180
agggtgttgt gagtgtaaat tagtgcgatg agtaggggaa gggagcctac tagggtgtag
                                                                         240
aataggaagt atgtgcctgc gttcaggcgt tctggctggt tgcctcatcg ggtgatgata
                                                                         300
gccaaggtgg ggataagtgt ggtttcgaag aagatataaa atatgattag ttctgtggct
                                                                         360
                                                                         420
gtgaatgtta taattaagga gatttgtaag ggagattagt atanagaggt anagtttttt
tcgtgatagt ggntcactgg ataantggcc gttggctttg ccatgattgt gaggggtagg
                                                                         480
agtcaagtag ttagtattan gangggggtt nttaggggtc cnaggaaang ttggggaana
                                                                         540
ctaaannggt gtngtnattn gtaaaaaata nnnnanggat n
                                                                         581
      <210> 463
      <211> 574
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (574)
      <223> n = A, T, C \text{ or } G
      <400> 463
actgtgtggc gccttattct aggcacttgt tgggcagaat gtcacacctg ccgatgaaac
                                                                          60
tectgegtaa gaagategag aageggaace teaaattgeg geageggaac etaaagttte
                                                                         120
agggggcctc aaatctgacc ctatcggaaa ctcaaaatgg agatgtatct gaagaaacaa
                                                                         180
tgggaagtag aaaggttaaa aaatcaaaac aaaagcccat gaatgtgggc ttatcagaaa
                                                                         240
ctcaaaatgg aggcatgtct caaqaaqcaq tqqqaaatat aaaaqttaca aaqtctcccc
                                                                         300
agaaatccac tgtattaagc aatggagaag cagcaatgca gtcttccaat tcagaaccaa
                                                                         360
aaaaaaaaa naaaaaaaag tactttttt ttttnnnnnt ttttttttt taggtaatgg
                                                                         420
gtgttgagct tgaacgcttt cttaattggn ggctgctttt angcctctat gggtgttaaa
                                                                         480
ttttttactc tcttacaaqq tttttcctaa qtccaaanac tqtccttttq qctacaqtta
                                                                         540
aatttccagg ggattaaagg gttttgggcn aatt
                                                                         574
      <210> 464
      <211> 580
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (580)
      <223> n = A, T, C \text{ or } G
      <400> 464
ggtacctagt aagctctccc tcctcccacc ctccaccctc aaggaggccc cagtgtcagt
                                                                          60
```

```
120
tgttcccctc tgggtccatg agttcttatc atttagctcc cacttataag caagaacatg
                                                                            180
cagtatttgg ttttctgttc ctgccttagt ttgctaagga taacggcctc cagctccatc
                                                                            240
cagtteetge aaaggacatg atcetgttet ttetatgget gtatagtatt ceatggtgta
                                                                            300
tatttaccae attgtettta tecagtetgt cattgatggg ettttgggtt gattagtage
                                                                            360
tttttgaatg gtaacttttc tacagaagta cgcggggctt ttttttttgc tgtaggcccg
                                                                            420
ggtggttgct gccgaaatgg gcangttcat gaaacctggg aaggtggtgc ttgtcctgct
                                                                            480
ggacgctact neggacgcaa agetgtcate gtgaaagaac attgatgatg geacettana
                                                                            540
cgccctacag ccatgctctg gtggctggaa ttgaccgcta cncccgaaag tgacagctgn
catgggcaag aagaagatcg ccagagatca aagataaaan
                                                                            580
      <210> 465
      <211> 578
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (578)
      <223> n = A, T, C or G
      <400> 465
ggtacttttt ttttttttt ttttttttt ttctacatca ctttanaata tttattgtat
                                                                             60
                                                                            120
teettaatge atticttaae atgiatagea etetttaate aagaatataa agicatetae
ttagaatcac attatettaa agatgeatae tggaatgata agtttgaaga tgtaactate
                                                                            180
                                                                            240
aacaattott ttoaaaatoa tatoaatata ttactotoat ggaacttgca cattotaaga
                                                                            300
agggtcattt tttcccccca gtaccaatat tacattattt gacagggata ataaaatgag
cagagactgg aaatcacaga caataacatt gettteteaa ttaacagaaa ggatteataa
                                                                            360
catatteett aaeggtagat gtgatttgta gagaatgtgg aaaagaacta ttgagaagte
                                                                            420
cacctgctgc ccaaactgag gcacattagg gtggttgtgg gangagttat atttgagggt ccatttttcc ttagggtta aaagcatgtc cnggttggng gtnatttgcc attaagtctn
                                                                            480
                                                                            540
                                                                            578
ttttcaaata aaagaattag gggagaaagt ttggaaaa
      <210> 466
      <211> 546
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(546)
      <223> n = A, T, C \text{ or } G
      <400> 466
accaatacca ccaattttgt agacatcctg gagaggcagg cgcaagggct tgtcagttgg
                                                                             60
acgagttggt ggtaggatgc agtccagagc ctcaagcagc gtggttccac tggcattgcc
                                                                            120
atcettacgg gtgactttcc atcecttgaa ccaaggcatg ttagcacttg getecagcat gttgtcacca ttccaaccag aaattggcac aaatgctact gtgtcggggt tgtagccaat
                                                                            180
                                                                            240
tttcttaatg taagtgctga cttccttaac aatttcctca tatctcttct ggctgtaggg
                                                                            300
tggctcagtg gaatccattt tgttaacacc gacaattagt tgtttcacac ccagtgtgta
                                                                            360
agccagaang gcatgctctc gggtctgccc attcttggag ataccagctt caaattcacc
                                                                            420
aacaccagca gcaacaatca ggacagnaca gtcggnctga gatgtccctg taatcatgtt
                                                                            480
ttgataaaag tetetgtgte etggggeate aatgatagte acatagtace teggeegega
                                                                            540
ncacgc
                                                                            546
```

```
<210> 467
      <211> 445
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (445)
      \langle 223 \rangle n = A,T,C or G
      <400> 467
acctaaaacc cgaagaacct tctgtaagaa gtgtggcaag catcagcctc acaaagtgac
                                                                         60
acagtataag aagggcaagg attetttgta tgeecaggga aggaggeget atgateggaa
                                                                        120
gcagagtggc tatggtgggc agacaaagcc aattttccgg aagaaggcta agaccacaaa
                                                                        180
gaagattgtg ctaaggctgg aatgtgttga gcctaactgc agatccaaga ggatgctggc
                                                                        240
tattaagaga tgcaagcatt ttgaactggg aggagataag aagagaaagg gccaagtgat
                                                                        300
ccagttctaa actttgggat atttttcttc aattttgaag agaaaatggt gaaccataga
                                                                        360
aaagttaccc gagggaaaat aaatacagtg atattccaaa aaaaaaaann nnnnnaaaaa
                                                                        420
aaagtnettg geegggaeee eetaa
                                                                        445
      <210> 468
      <211> 566
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (566)
      <223> n = A,T,C or G
      <400> 468
actgtgtggc gccttattct aggcacttgt tgggcagaat gtcacacctg ccgatgaaac
                                                                         60
tectgegtaa gaagategag aageggaace teaaattgeg geageggaac etaaagttte
                                                                        120
agggggcctc aaatctgacc ctatcggaaa ctcaaaatgg agatgtatct gaagaaacaa
                                                                        180
tgggaagtag aaaggttaaa aaatcaaaac aaaagcccat gaatgtgggc ttatcagaaa
                                                                        240
ctcaaaatgg aggcatgtct caagaagcag tgggaaatat aaaagttaca aagtctcccc
                                                                        300
agaaatccac tgtattaagc aatggagaag cagcaatgca gtcttccaat tcagaaccaa
                                                                        360
aaaaaaaaa nnaaaaaaag tactttttt tntnnnnnn ttttttttag gaatgggtgt
                                                                        420
tgaacttgac ctttcttaat gggggctggt tttaggctat atggngtaaa tttttctctt
                                                                        480
ttacaaggtt tttcctagng ncaaaaactg tcctttggac taccgtaaat tacaggggtt
                                                                        540
taaaggttnt ggggcaatta aanttn
                                                                        566
      <210> 469
      <211> 586
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (586)
      <223> n = A, T, C \text{ or } G
```

```
<400> 469
acgcgggata ggtttggtcc tagcctttct attagctctt agtaagatta cacatgcaag
                                                                    60
                                                                   120
catccccgtt ccagtgagtt caccctctaa atcaccacga tcaaaaaggga caagcatcaa
                                                                   180
gcacgcagca atgcagctca aaacgcttag cctagccaca cccccacggg aaacagcagt
                                                                   240
gattaacctt tagcaataaa cgaaagttta actaagctat actaacccca gggttggtca
atttcgtgcc agccaccgcg gtcacacgat taacccaagt caatagaagc cggcgtaaag
                                                                   300
agtgttttag atcacccct ccccaataaa gctaaaactc acctgagttg taaaaaactc
                                                                   360
cagttgacac aaaatagact acgaaagtgg ctttaacata tctgaacaca caatagctaa
                                                                   420
                                                                   480
gacccaaact qqqattaqat accccactat gcttagccct aaacctnaca gttaaatcaa
caaaactgct cgccagacac tcgagccaca gcttaaaact caaggacctg cgggcttcat
                                                                   540
atccctctag angacctgtc tgtaatcgat aaccccgatc aacctn
                                                                   586
     <210> 470
     <211> 487
     <212> DNA
     <213> Homo sapiens
     <220>
     <221> misc feature
     <222> (1)...(487)
     <223> n = A, T, C \text{ or } G
     <400> 470
                                                                    60
acggccaggg ctattggttg aatgagtagg ctgatggttt cgataataac tagtatgggg
                                                                   120
ataaggggtg taggtgtgcc ttgtggtaag aagtgggcta gggcattttt aatcttagag
180
                                                                   240
                                                                   300
                                                                   360
atcatttgtt ttgaggttag tttgattagt cattgttggg tggtgattaa tcngttgntg
                                                                   420
atgaaatatt tggaggtggg gatcaatana gggggaaata gaatgatcag tacctcgccc
                                                                   480
gegaceaege taagggeeaa teeacaeaet ggeggnegta etaatggate ceaaetegga
                                                                   487
ccagctt
     <210> 471
     <211> 488
     <212> DNA
     <213> Homo sapiens
     <220>
     <221> misc feature
     <222> (1)...(488)
     <223> n = A, T, C or G
      <400> 471
                                                                    60
actgcggcgg gtaggcctag gattgtgggg gcaatgaatg aagcgaacag attttcgttc
attttggttc tcagggtttg ttataatttt ttattttat gggctttggt gagggaggta
                                                                   120
ggtggtagtt tgtgtttaat atttttagtt gggtgatgag gaatagtgta aggagtatgg
                                                                   180
                                                                   240
gggtaattat ggtgggccat acggtagtat ttagttgggg cattcccgcg tacctatttg
                                                                   300
tatttttggt agagacaggg ttttgccatg ttggccagga tggtcttgaa ctactgacct
                                                                   360
caggtgatcc tcacgccttt atctcccaaa gtgctgcgat tacaggcatg aggcaccact
                                                                   420
cctggccaca ttcttatatt taaaaaaaaa gcacaactct attgtctact ggtgttcttt
480
                                                                   488
naccacnc
```

11.4

```
<210> 472
      <211> 586
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (586)
      \langle 223 \rangle n = A,T,C or G
      <400> 472
                                                                            60
ggtacttgat gccctccaag caattaaaac caagggcaaa cgagccccat tcacaaattt
tgacccctct actctccttc cttcatccct ggatttctgg acctaccctg gctctctgac
                                                                           120
tcatcctcct ctttatgaga gtgtaacttg gatcatctgt aaggagagca tcagtgtcag ctcagagcag ctggcacaat tcagcagcct tctatcaaat gttgaaggtg ataacgctgt
                                                                           180
                                                                           240
coccatgoag cacaacaacc goccaaccca acctotgaag ggcagaacag tgagagotto
                                                                           300
attttgatga ttctqaqaaq aaacttqtcc ttcctcaaqa acacagccct gcttctgaca
                                                                           360
taatccagta aaataataat ttttaagaaa taaatttatt tcaatattag caaagacagc
                                                                           420
atgeetteaa ateaatetgt aaaactaaga aaettaaatt ttagttetta etgettaate
                                                                           480
                                                                           540
aaataataat tagtaagcta gcaaatagta atctgtaagc ataagcttat gcttaaatca
                                                                           586
gtttagtttg aggaatcttt aaaattacca ctaantgatt gnatgg
      <210> 473
      <211> 575
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (575)
      <223> n = A, T, C or G
      <400> 473
                                                                            60
ggtacaaagg ggaaagggtg catgccaact atcgaattat aggatatgta aaaaatataa
gtcaagaaaa tgccccaggg cccgcacaca acggtcgaga gacaatatac cccaatggaa
                                                                           120
                                                                           180
ccctgctgat ccagaacgtc acccacaatg acgcaggaat ctatacccta cacgttataa
                                                                           240
aagaaaatct tgtgaatgaa gaagtaacca gacaattcta cgtattctcg gagccaccca
agccctccat caccagcaac aacttcaatc cggtggagaa caaagatatt gtggttttaa
                                                                           300
cctgtcaacc tgagactcag aacacaacct acctgtggtg ggtaaacaat cagagcctcc
                                                                           360
tggtcagtcc caggctgctg ctctccactg acaacaggac cctcgttcta ctcacgccca
                                                                           420
                                                                           480
aagaatgaca taggacccta tgaatgtgaa atacagaacc cagtgggtgc cacccgcant
gcccantcac cctgaatgtc cgtatgagtc aatcctgccg gcggccgttc naanggcgaa
                                                                           540
ttccacacac tggcggccgt ctaatggatc cactc
                                                                           575
       <210> 474
       <211> 515
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (515)
```

```
<223> n = A, T, C \text{ or } G
      <400> 474
                                                                           60
ggtacgtggg ggactcaact gaaatcatgg cgtttgacag cacttggaag gtagaccgga
                                                                          120
gtgaaaacta tgacaagttc atggaaaaaa tgggtgttaa tatagtgaaa aggaagcttg
                                                                          180
cageteatga caatttgaag etgacaatta cacaagaagg aaataaatte acagteaaag
aatcaagcgc ttttcgaaac attgaagttg tttttgaact tggtgtcacc tttaattaca
                                                                          240
acctagcaga cggaactgaa ctcaggggga cctggagcct tgagggaaat aaacttattg
                                                                          300
gaaaattcaa acggacagac aatggaaacg aactgaatac tgtccgagaa attataggtg atgaactagt ccagacttat gtgtatgaag gagtagaagc caaaaggatc tttaaaaagg
                                                                          360
                                                                          420
attgaccatt attcttggcg cacagtccaa aatncaaatt ggccagaaga tctatattgn
                                                                          480
                                                                          515
acctgcccgg gcggccgttc gaaaggccaa ttcca
      <210> 475
      <211> 580
      <212> DNA
      <213> Homo sapiens
      <221> misc feature
      <222> (1) ... (580)
      <223> n = A,T,C or G
      <400> 475
acaaagatct gacatgtcac ccagggaccc atttcaccca ctgctctgtt tggccgccag
                                                                           60
tettttgtet etetetteag caatggtgag geggatacee ttteeteggg gaagagaaat
                                                                          120
                                                                          180
ccatggtttg ttgcccttgc caataacaaa aatgttggaa agtcgagtgg caaagctgtt
gccattggca tettteaegt gaaceaegte aaaagateea gggtgeetet etetgttggt
                                                                          240
gatcacacca attettecta ggttagcace tecagteace atacacaggt taccagtgte
                                                                          300
gaacttgatg aaatcagtaa tottgocagt ototaaatca atotgaatgg tatcattcac
                                                                          360
cttgatgagg ggatcggggt agcggatggt gcgggcatca tgagtcacca gatgagggat
                                                                          420
                                                                          480
tccttttgtg ccccaaagat ctttctnact ttgacaactt gaccttggnc gcgaccaccc
                                                                          540
taaggegaat teacceactg geggeegtet aatggateen neteggneea acetggntat
                                                                          580
atggentaan tnntcenggn naaatntnte ecencaatee
      <210> 476
      <211> 593
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (593)
      <223> n = A,T,C or G
      <400> 476
ggtactatgt gggacagtat tttgcaaata caagaagagc tcagggcagc tgtggagctg
                                                                           60
gatggtetge etggeaggee tetgtgeagt etgeetgete atectgteee etttttgggg
                                                                          120
cttgatecte tteteggigt catgetteet catgtatact tacttatetg gecaagaatt
                                                                          180
                                                                          240
gttacctgtg gatcagaagg cagtcctggt gacaggtgtg attgcgggct tggccatgct
                                                                          300
ttgtgcaagt atctggatga gctgggcttc acggtatttg ccggagtttt gaatgaaaat
ggcccaggag ctgaggaatt gcgaagaacc tgctctccgc gcctctcggt gctccaaatg
                                                                          360
gacatcacga accagtgcag ataaaagatg cttacagcaa ggttgcaaca atgctgcagg
                                                                          420
```

```
480
acaaaaqact qtqqqctqtq atcaacaatq ctnqqqtqct tqqcttttcc actqatqqqq
                                                                           540
agettnttnt tatgatgact acnaacaatc ntggccgnga acttttttga actgngaggg
acaaaacgtt tttccttttt taaaaaancc aagggnggtg gnnaattncn nnt
                                                                           593
      <210> 477
      <211> 595
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(595)
      \langle 223 \rangle n = A,T,C or G
      <400> 477
actacaaggt ttagcatttg ctctgctggt cgacattccc ccagtctatg ggttgtatgc
                                                                            60
atcettttte ccagecataa tetacetttt etteggeact tecagacaca tatcegtggg
                                                                           120
teegttteeg attetgagta tgatggtggg actageagtt teaggageag ttteaaaage
                                                                           180
agteceagat egeaatgeaa etaetttggg attgeetaae aactegaata attetteaet
                                                                           240
actggatgac gagagggtga gggtggcggc ggcggcatca gtcacagtgc tttctggaat
                                                                           300
catccagttg gcttttggga ttctgcggat tggatttgta gtgatatacc tgtctgagtc
                                                                           360
cctcatcagt ggcttcacta ctgctgctgc tgttcatgtt tttggnttcc caactcaaat
                                                                           420
tcatttttca agtgacagtc ccgtcacaca ctgatncagt ttnaatttta aaagtacctc
                                                                           480
ggccgcganc accctaaggc gaatttnaac ccactngcgg ccgttctant ggatccaact ngnnncaaac ttngngaata ngggcataac ngntcctggg gaaatnnttc ccnct
                                                                           540
                                                                           595
      <210> 478
      <211> 420
      <212> DNA
      <213 > Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (420)
      <223> n = A,T,C or G
      <400> 478
ggtacacagt atgtataaca atgcatacta tggtgtggag ttaattccaa ttaccatatt
                                                                            60
ttatatttat tggtcacaac agcatacatt ttatgctcca aaatacatgg atctgacaaa
                                                                           120
atggttacat ttaatgttct tttaaagaaa gatgaactaa atttaagaag aattggtttt
                                                                           180
tectaatate teatttteaa attactgata caaatttgee agagaaacaa ttacatgttt
                                                                           240
tacctaacat caaataatct ccagtticta agacagatgc atticttqtt caatticcaa
                                                                           300
aagtaaataa aggettteta aetgaaaaca titgeateee tageteteta aagtaattaa
                                                                          360
aaagaaaatt acaaaaaatg acctctaagc ttctgaacag cccacttant tacataaagt
                                                                           420
      <210> 479
      <211> 602
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (602)
```

```
<223> n = A, T, C or G
      <400> 479
ggtacctagt cagatggtag acgagetgte tgetgeegea ggageacete tatacaggae
                                                                           60
ttagaagtag tatgttattc ctggttaagc aggcattgct ttgccctgga gcagctattt
                                                                          120
taagccatct cagattetgt ctaaaggggt titttgggaa gacgttttct ttatcgccct
                                                                          180
gagaagatet accccaggga gaatetgaga catettgeet actittett attagettte
                                                                          240
tecteateea titetittat acctitecti titiggggagt igitatgeea igatititigg
                                                                          300
tatttatgta aaaggattat tactaattct atttctctat gtttattcta gttaaggaaa
                                                                          360
tgttgagggc aagccaccaa attacctang ctgaggttag agagattggc cagcaaaaac
                                                                          420
tgtgggaaga tgaactttgt cattatgatt tcattatcac algattatag aaggctgtct
                                                                          480
taatgcaaaa aacatactta catttnanac atattccaan gggatctcnc attttgtaaa
                                                                          540
aagttgacta ttactggagt aaaccctgtt ttccctaant ttaacttttt ttgggaaatt
                                                                          600
                                                                          602
      <210> 480
      <211> 600
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (600)
      <223> n = A, T, C or G
      <400> 480
ggtacttttt ttttttttt ttttttttc ggtttgaggg ggaatgctgg anattgtaat
                                                                           60
gggtatggan acatgtcata taagtaatgc tagggtgagt ggtaggaagt tttttcatag
                                                                          120
gaggtgtatg agttggtcgt agcggaatcg ggggtatgct gttcgaattc ataaaaacag ggaggttana agtagggtct tggtgacaaa atatgttgtg taaagttcag ggganagtgc
                                                                          180
                                                                          240
gtcatatgtt gttcctagga aaattgtagt ggtgagggtg tttattataa taatgtttgt
                                                                          300
gtattegget atgaaaaata gggegaaggg geetgeggeg tatteeatgt tgaageetga
                                                                         360
gactagtteg gacteceett eggeaaggte caaaggggtt eeggttggte tettetaqtg
                                                                          420
tggagataaa tcatattatg gccnagggtc atgatggcag gagtaatcaa aggggtcntt
                                                                          480
tgttttgaaa aagggnggan aggttaaagg ancccctttt tataatggtg atantaaaaa
                                                                         540
gatgettggg ggactenttt aaaatgttgg etettettee angeneeeac aggegtattt
                                                                          600
      <210> 481
      <211> 594
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (594)
      <223> n = A,T,C or G
      <400> 481
cgaggtacgg ccagggctat tggttgaatg agtaggctga tggtttcgat aataactagt
                                                                          60
atggggataa ggggtgtagg tgtgccttgt ggtaagaagt gggctagggc atttttaatc
                                                                         120
ttagagegaa agectataat caetgegeee geteataagg ggatggeeat ggetaggttt
                                                                         180
atagatagtt gggtggttgg tgtaaatgag tgaggcagga gtccgaggag gttagttgtg
                                                                         240
gcaataaaaa tgattaagga tactagtata agagatcagg ttcgtccttt agtgttgngt
                                                                         300
```

```
atggttatca tttgttttga ggttagtttg attagtcatt gttgggtggt gattantccg
                                                                       360
                                                                       420
ttqttqatqa qatatttgga ggtggggatc aatagagggg gaaatagaat gatcagtacc
tgccenggeg gnegetegaa anggegaatt ceaceaeat ggegggegnt ctaatggatn
                                                                       480
cgaccongtc ccaacttgcg taatcatggc atacttgttn ctggtgaaat ggtatccctc
                                                                       540
                                                                       594
acaattccca cacatacaac ccgaacctaa atgtaaanct gggggcctat natn
      <210> 482
      <211> 600
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(600)
      <223> n = A,T,C or G
      <400> 482
accatgaaat acatatattt cataaggttc agttacaaaa tggattgttt caaatggcaa
                                                                        60
tttcttacac taacctgatt atgaaaaaaa gaagtctgta tcatctgctt ccaagtctgt
                                                                       120
tatqtccaaa tatattttaa ttatqcattt attttqctac ttttataaat attagagatt
                                                                       180
tcaccttaaa ttatttttgt aactagttct agaacatgtt ttccaattat tatttttcta
                                                                       240
atggagacat ataattgacc tatgtttatg catatatgtt ctctacacag tgaaactttt
                                                                       300
tttaaaaaga atagtaaaga aaatgcggaa gctctggctc tccaaggcaa agtcaaaaaa
                                                                       360
aaaaaaaag cggggggaa tgcgaggaac attttattac acctnctgat tttcctcctt
                                                                       420
gagntttatt ttctcccctt ggntatttgt taatgctaga aactgnattc ctaanaaagc
                                                                       480
atacctettt caggngagen tgataattgg gaanaatttt gtteetttag tntgaacatt
                                                                       540
ttattaagaa gngatteeta ataaaganae aangggetnt ttaattnttt gggggnngga
                                                                       600
      <210> 483
      <211> 605
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (605)
      <223> n = A,T,C or G
      <400> 483
acagaacatc qtcaqcacta qcacaqttta caqaacctca caqacccaaa qgaacatcaa
                                                                        60
                                                                       120
taggcaaagc gactacagga ggcgtgtgtc cgcgtgggcg aggtaaagag ggtcagtatt
                                                                       180
ggtcaagtga cagtgtcggt aatctggcaa gacagtgatg ttaagaaggt tcatagttta
agaattatet aaaatattit aaaaactata aagetgeaac acatgattit tacacetagt
                                                                       240
tactagaaaa ctaaggaaag cacttattag ctctgaataa agtaacatgg aaagcacttt
                                                                       300
tactaatcga caaaaaaacc ttctaatgca ttatcagaaa gattttataa tacaaggagg
                                                                       360
                                                                       420
catattgctc agtcagaagg ggttctataa gaaaagcact tactaagtta gcgactaaca
gaacaaceng tttaaagatg aattaaatgc cccatttggg gangcatggc aggtgttaag
                                                                       480
anaaangaaa agentaagaa aacatttnet ggttatanea aacetttntt tnttatetae
                                                                       540
tgnatttgac aaaaattaac cntttaaaqt tttaccengq cacttnnttc nttgtcctcg
                                                                       600
gcccg
                                                                       605
      <210> 484
      <211> 591
```

```
<212> DNA
      <213> Homo sapiens
      <221> misc feature
      <222> (1) ... (591)
      <223> n = A.T.C or G
      <400> 484
ggtacgcggg tggggagacc ctggggtagc agccactgac ctcacacctg gaggaagctg
                                                                        60
tgtgaccgat tcatgagett atgeetgaag acagageaag cacteecege accaegaega
                                                                       120
tgacgttcac ttgttttgtg tttttcgatc tcttcaacgc cttgacctgc cgctctcaga
                                                                       180
ccaagetgat atttgagate ggetttetea ggaaceacat gtteetetae teegteetgg
                                                                       240
ggtccatcct ggggcagctg gcggtcattt acatccccc gctgcagagg gtcttccaga
                                                                       300
eggagaacet gggagegett gatttgetgt ttttaactgg attggeetea teegtettea
                                                                       360
ttttgtcaga gctcctcaaa ctatgtgaaa aatactgttg cagccccaaa gagagtccag
                                                                       420
atgcaccetg aaagatgtgt agtggacege actteegegg nacetteeta atnattteaa
                                                                       480
ctgggtgnga ctgtggccct gccctgtttc ttcttagggg agactttang anggcgagcn
                                                                       540
tcataccgga tagttttctt taggaaactn aggaaccttg gctcaggacc a
                                                                       591
      <210> 485
      <211> 605
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(605)
      <223> n = A,T,C or G
      <400> 485
ggtacgcggg gatataaagg gagagagcaa gcagcgagtc ttgaagctct gttnggtgct
                                                                        60
tnggatccat ttccatcggn ccttacagcc gctcgtcaga ctccancagc caanatggtg
                                                                       120
aancagatcg agagcaagac tgcttttcan gaagccttgg acgctgcang tgataaactt
                                                                       180
gtagnagttg acttctcagc cacgtggtgt gggccttgca aaatgatcaa gcctttcttt
                                                                       240
cattccctct ctgaaaagta ttccaacgtg atattccttg aagtagatgt ggatgactgt
                                                                       300
caggatgttg cttcagagtg tgaagtcaaa tgcatgccaa cattccagtt ttttaagaag
                                                                       360
ggacaaaagg tgggtgaatt ttctggagcc aataaggaaa agctttgnag ccnccattaa
                                                                       420
tgaatgagtc taatcatqtt ttctqaaaac ataacccaqc catttqqcta tttaaaactt
                                                                       480
gnaanttttt nagntacena aatttaaagt etgaagaeat aacceggtge catttgegtg
                                                                       540
                                                                       600
acaatnaaaa attatgccaa cactttttna anaanganan nnntttcctn gggaaatngt
anccc
                                                                       605
      <210> 486
      <211> 319
      <212> DNA
      <213> Homo sapiens
      <400> 486
ggtaccagtt gtagccataa agattctggg actcattatg gactactaga aggacctcct
                                                                        60
tecettetge gacattgaac ggcacgacat caatattggt etgggcactg titggcaggt
                                                                       120
tccagaaggt taaaagcgag gctgtgagca ggagtccctg ccagggaatg cacactctgt
                                                                       180
                                                                       240
atggacagge tgaaggggac eccatqqtet etgetgeetg ettgteetet gtggagaaga
```

```
gettgggete caggaactet ettgtcaggg etgetgtgae tgtcagetet getgteette
                                                                          300
ctacctctgt gtcccqcqt
                                                                          319
      <210> 487
      <211> 586
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (586)
      <223> n = A,T,C or G
      <400> 487
acgcgggagc tgagtgtccc gcggggcccg aagcgtttac tttgaaaaaa ttagagtgtt
                                                                           60
caaagcaggc ccgagccgcc tggataccgc agctaggaat aatggaatag gaccgcggtt
                                                                          120
ctattttgtt ggttttcgga actgaggcca tgattaagag ggacggccgg gggcattcgt
                                                                          180
attgcgccgc tagaggttaa attettggac cggcgcaaga cggaccanag cgaaagcatt
                                                                           240
tgccaagaat gttttcatta atcaagaacg aaagtcggag gttcgaagac gatcagatac
                                                                          300
cgtcgtagtt ccgaccataa acgatgccga ccggcgatgc ggcggcgtta ttccatqacc
                                                                          360
cgccgggcag ctttcnggaa accaaagtct ttgggttncc gggggagtat ngttcnaaaa
                                                                          420
aaaaaaaaa aaaaaaagt cctnggccgg gancccctta nggngaaatt cagccactgg
                                                                          480
nggcgttctn atggatncna gctcggncca acntggcgta atatggcata cttgttcctg
                                                                          540
gngnaaatgt ttccctccaa attccccaaa tacgggcgga gcttaa
                                                                          586
      <210> 488
      <211> 487
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (487)
      <223> n = A,T,C or G
      <400> 488
acagetggtt ggacetatte atgeatette accageaget ggageatete caccettggt
                                                                           60
atttctggtg taaattactt gagetetgtg etttgaaace agtttgataa gteetttact
                                                                          120
aaggagetee tgaagggetg eeetggeeag ggageetega aletteagte leteagagae
                                                                          180
cacagetggg gttataagtt tatagttggg aactteetta cagagtttat cataggtage
                                                                          240
tttgtcaaac aagactaagt tattgagctt gtcccgaact ttgcctttgg accacttctt
                                                                          300
ctttttggcc ttgccccgg atttgttcac tgggtctttg tctttcttgg ccgactttcc agcgtccttc ttcttcttgt cgtccttaag cggcattgcg aanctcggag aataagcaac
                                                                          360
                                                                          420
aaacaccgca cctcgtcnaa gatgtcggac aaaaaaaaggc cccgcgtacc ttnggccgcg
                                                                          480
ancacno
                                                                          487
      <210> 489
      <211> 589
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
```

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```
<222> (1)...(589)
       <223> n = A, T, C or G
      <400> 489
acgcggggtc tctcctcagg cagcagcaac gcggaggaaa cgggagtgaa cggagagcgt
                                                                            60
agtgaccatc atgagectee teaacaagee caagagtgag atgaceccag aggagetgea
                                                                           120
gaagcgagag gaggaggaat ttaacaccgg tccactctct gtgctcacac agtcagtcaa
                                                                           180
gaacaatacc caagtgctca tcaactgccg caacaataag aaactcctgg gccgcgtgaa
                                                                           240
ggccttcgat aggcactgca acatggtgct ggagaacgtg aaggagatgt ggactgaggt
                                                                           300
acaaagatta aattaagaca cggtaaattg actaaatatt tggtttttat ataaataaag
                                                                           360
gtcataacca caccgttgac atgtaatact gttataatac aacagttaaa ctttgtgagt
                                                                           420
ctcaacagaa gtcatctgta gttnaacagg aaacaaaagt tgaaaaaaaa catgttnaaa
                                                                           480
caaaactctg ggactaacag gtcgggattg taagtacaac caacatattc ctcacttctg
                                                                           540
ggtntttcaa gtttacagta cttggccgga cccccttang ggnattcac
                                                                           589
      <210> 490
      <211> 591
      <212> DNA
      <213> Homo sapiens
      <221> misc feature
      <222> (1)...(591)
      <223> n = A, T, C \text{ or } G
      <400> 490
ggtaccggga tagtttttgc agggttttat tttataaaat ccaagcgcgc tgttgattgt
gttttccttg ttttcagccc cccgactcca gcccgcagca catttccgct gtccgtcagt
                                                                          120
aattgtgtcc tctctttatg cttgcttggg gaatgttgtt ttctgactag gctgatcatt
                                                                          180
atctaaagaa tctaattctg ttgattttta aaacttttag gaccataaac gttgtgttca tatatggaca tggaaatatt tatataattt tatagaaaat aaccttttag atggtcaaag
                                                                          240
                                                                          300
tgtaaggagt tttttttgtc agataatcat ttctacttca aaaacatttc atgcaatatt
                                                                          360
agaataaagt teetgteatt eetetnnnan aaaaannnnn nnnnnnanna nnnnnnnnn
                                                                          420
nggaanannn nnnnnnnnn aaaaaagtac ctgcccnggc ggccgttcaa aaggcgaatt
                                                                          480
ccacccactg geggeegtte taatggatee ancteggace aacetggnga aacatggeat
                                                                          540
acctgttcct ggngaaatgg tntcccttac aattcccaca aataaaaccg g
                                                                          591
      <210> 491
      <211> 583
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (583)
      \langle 223 \rangle n = A,T,C or G
      <400> 491
ggtacatata aatccttttg gtgtcacttg tagcaagcct tgcttctgca gttttcggat
                                                                           60
tttcctcaaa getttgttge gettgegtag aattegaagt ggactaaage caacageate
                                                                          120
gataagtttc cgcctaaaga aaccaatgtt tgcaaagtag ataggagatg gacatctgaa
                                                                          180
aattttcact ccttctggct catacatatc ataataatct tttttattct tatagatgtt
                                                                          240
ggttcttcca atattagcca gcgtgctgca ttttggaaat tgggtcctga acacgatggt
```

300

```
tagcagttga aatgccacac tagctgccag gcctaacccg agtcccagga caatggtgaa
                                                                           360
agatgaaagg catgaaccca aataaacaat catatttggn cnttccccca atctgctatt
                                                                           420
ttaaccaact gcatcaacat tcctttaagt tccaatgcta aactggcang acnggenttt
                                                                           480
gtagaagngc cangaaaaat cagngcttga cgacaatcac accatgatgn nccataancc
                                                                           540
acaatctggg nttggctcnn ggcctctgaa cnnngactgg nag
                                                                           583
      <210> 492
      <211> 597
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (597)
      <223> n = A, T, C or G
      <400> 492
acgcgggggg tggcacggag gaaccaggag cgtgccctgc gcaccgtctg gagctccgga
                                                                           60
gatgacaagg agcagctggt gaagaacaca tatgtcctgt gaccgccctg tcgccaagag
                                                                          120
gactggggaa gggaggggag actatgtgtg agcttttttt aaatagaggg attgactcgg
                                                                          180
atttgagtga tcattagggc tgaggtctgt ttctctggga ggtaggacgg ctgcttcctg
                                                                          240
gtctggcang gatgggtttg ctttggaaat cctctangag gctcctcctc gcatggcctg
                                                                          300
cagnotggca acaaccocga gttgtttcct cgctgatcga tttctttcct ncaggtagag
                                                                          360
ttttctttgc ttatgttgaa ttccattgcc tttttctcat cacaaaaaat gatgttggga
                                                                          420
atcgnntctt ttgtttggct gaattatggg ntttttaant ataaaccaaa nttttttatt
                                                                          480
aacattetta aanaagggaa agtnnaatgt nenttggnee enaceneget aanggenaat
                                                                          540
ttcanecent ggnggeegtn nttnnggate ennnenngnn ecaannntgg nntantn
                                                                          597
      <210> 493
      <211> 591
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (591)
      \langle 223 \rangle n = A,T,C or G
      <400> 493
acggatgcta cttgtccaat gatggtaaaa gggtagctta ctggttgtcc tccgattcag
                                                                           60
gttagaatga ggaggtctgc ggctaggagt caataaagtg attggcttag tgggcgaaat
                                                                          120
attatgettt gttgtttgga tatatggagg atggggatta ttgctaggat gaggatggat
                                                                          180
agtaataggg caaggacgc tectagtttg ttagggacgg ateggagaat tgtgtaggeg
                                                                          240
aataggaaat atcattcggg cttgatgtgg ggaggggtgt ttaaggggtt ggctagggta
                                                                          300
taattgtctg ggtcgcctag gaggtctggt gagaatagtg ttaatgtcat taaggagaga
                                                                          360
aggaagagaa gtaagccgag ggcgtctttg attgtgtagt aagggtggaa ggtgatttta tcggaatggg aggtgattcc taagggggtg gttgatcccg tttcctgcca agaataagaa
                                                                          420
                                                                          480
gtggaatget getagggetg cattaatgaa ggccaagatg aaatgaaagg taaanaaten
                                                                          540
ngtgangggg gactgctact gatancetet caaatcatga ataggntgte c
                                                                          591
      <210> 494
      <211> 374
      <212> DNA
```

```
<213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (374)
       <223> n = A, T, C or G
       <400> 494
ggtacttttt ttttttttt ttttttttt ttttttagnt catgtctttt attaactcat
                                                                               60
acagttactt gtcttctggt ttgttgaaac agtaagtcan acaacatttg ccacaataat
                                                                              120
gtotgtoaaa gtgaottgco ataaacacco cagcaccaca ttoatcanaa gggcaototo
                                                                              180
gacgaaggcg actaattttg ccattctcat ccaccttata atatttcagg acagccagct
                                                                              240
taaccttctt tctcttgtgc ttattcttct tgggagnggt gtaagacttc ttcttccttt
                                                                              300
tcttagcacc accacgaagt ctcaacacaa gatgaagagt agactccttt tgaatattgt
                                                                              360
aagtcagaca aagt
                                                                              374
       <210> 495
       <211> 597
       <212> DNA
       <213> Homo sapiens
       <220>
      <221> misc_feature
      <222> (1) ... (597)
      <223> n = A,T,C or G
       <400> 495
actgggagaa ggtgctgacg ccgacgaagt ggtggatggg cttcccgctg caggtgaacc
                                                                               60
tcctggtgcc atcctgcagg gtcccccgag gattgcctag atcattttc aagcagtagt tgctttctgg gttttacaa attctgcatt ttccacactg aggagtaaag agcgggatga ctttatcacc tggtttgact gtagtcaccc cttctccaac actttccacg atgccggctg
                                                                              120
                                                                              180
                                                                              240
cctcatggcc taaaatcaca ggaaggggg tcaccaggtt gccactaacc acatgctcat
                                                                              300
ctgaacgaca gattcctgca gccaccatct taatgcgaac ttcatgagcc ttaggaggtg
                                                                              360
caacetetae etecteaatg gaaaagggtt tetttaacte ceatageaca actgetttge
                                                                              420
atttgattac ctgtaaactc agctacttgt gaaggctgag gcanganaat actttgaacc
                                                                              480
ccggaaggca aaggttgcaa tgagccnana acaccattgn acttccanct gggcaatana
                                                                              540
aaaaaactca tttttcctgc tggctcaaat gatctgcttc ttgcaaacaa gagntgn
                                                                              597
      <210> 496
      <211> 604
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (604)
      \langle 223 \rangle n = A,T,C or G
      <400> 496
ggacgcgggt gctgactgca tagctctttt tcttgagagg ctctccattt tgattcagaa
                                                                               60
agttagcata tttattacca atgaatttga aaccagggct ttttttttt tttgggtgat
                                                                              120
gtaaaaccaa ctccctgcca ccaaaataat taaaatagtc acatttatct ttattaggta
                                                                              180
atcacttctt aattatatgt tcatactcta agtatcaaaa tcttccaatt atcatgctca
                                                                              240
```

```
cctgaaagag gtatgctctc ttaggaatac agtttctagc attaaacaaa taaacaaggg
                                                                       300
gagaaaataa aactcaagga gtgaaaatca ggaggtgtaa taaaatgttc ctcgcattcc
                                                                       360
occeegettt ttttttttt ttgactttge cttggaaage cagagettee egeattttet
                                                                       420
ttactattct ttttaaaaaa agtttcactg ngtaaaagaa catatttgcc taaacatang
                                                                       480
tcaattatat gtctccatta naaaaaaata attggnaaac attgtctana actagttcca
                                                                       540
aaataattaa gggggaaatc tntaatnttt ttaaagtgcc naaanaatgc ctaanttaaa
                                                                       600
                                                                       604
      <210> 497
      <211> 587
      <212> DNA
      <213> Homo sapiens
      <221> misc feature
      <222> (1)...(587)
      <223> n = A,T,C or G
      <400> 497
acattaatga aatgtttcca aagaaatact gaacaatata tactctagtt tgctgaggtt
                                                                        60
ccagctcgag ttcaaaccta attcttgtgc aataaaaatc agcatggatc ttagatgatc
                                                                       120
tagaatacac tgtgttttga aatccacagc tggtttcatt tttaaccatt atgaaaaacc
                                                                       180
agtacttttt ttttttttt ttttttttc nctnggacca taaattttta ttggcaggtc
                                                                       240
aggaaaaaag ccgggggtaa gggtcccttc cttcccatcc ctctacccan aanacaccct
                                                                       300
ccaaaggaca gcagaagccc cagagcctgc tgcctcagag gaccttggag gcagacaaat
                                                                       360
tgttgtagng atcttcctgt ccctcaanca ggctgcggta ggtggnaatc tnctgctcca
                                                                       420
gccgcgactt gatgtccatg aaccgctggt cctcggccgc gacaccctta nggcgaattn
                                                                       480
caccnactgg gnggcgttct agtggatccg actcggacca acctngcgna atcatggcan
                                                                       540
actggttnct gnnggaaatg gtttccctnc aattccccaa cataccn
                                                                       587
      <210> 498
      <211> 354
      <212> DNA
      <213> Homo sapiens
      <400> 498
acgcgggcaa taaagctaaa actcacctga gttgtaaaaa actccagttg acacaaaata
                                                                        60
gactacgaaa gtggctttaa catatctgaa cacacaatag ctaagaccca aactgggatt
                                                                       120
agatacccca ctatgcttag ccctaaacct caacagttaa atcaacaaaa ctgctcgcca
                                                                       180
gaacactacg agccacagct taaaactcaa aggacctggc ggtgcttcat atccctctag
                                                                       240
aggageetgt tetgtaateg ataaaceeg ateaacetea ceaectettg eteageetat
                                                                       300
ataccgccat cttcagcaaa ccctgatgaa ggctacaaag taagcgcaag tacc
                                                                       354
      <210> 499
      <211> 632
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (632)
      <223> n = A,T,C \text{ or } G
```

```
<400> 499
necgaggtac caactgcact cgttttggca ttgcagctaa atatcagttg gatcccactg
                                                                           60
 cttccatttc tgcaaaagtc aacaactcta gcttaattgg agtaggctat actcagactc
                                                                          120
 tgaggcctgg tgtgaagctt acactctctg ctctggtaga tgggaagagc attaatgctg
                                                                          180
gaggccacaa ggntgggctc gccctggagt tggaggctta atccanctga aaagaaacct
                                                                          240
 ttgggaatgg atatcaaaag aattggcctt aatatatttc cattgngacc agcagcaggc
                                                                          300
tttttttccc ccagaagatg atcaaaacaa aaggatgatc tcaacaagaa ctgtatttta
                                                                          360
aagtatttaa ganagtottt ggtaactngg ttotaagtng gtatotaatt acccaatgot
                                                                          420
geagteetge agtecetatt cattanttaa atgtatttaa etggtaaatg ecetnecene
                                                                          480
cataatgaaa taganccttt ttgaaaaccc aaaaaaaaaa aaaaaaaaa aaaaaagtcc
                                                                          540
ctgcccggcc ggccctcaaa nggngaattc cannecctgg gggccgtact aanggatcen
                                                                          600
cccggnccaa cttggggaat atgggntant gn
                                                                          632
       <210> 500
       <211> 619
       <212> DNA
       <213> Homo sapiens
       <220>
      <221> misc feature
       <222> (1) ... (619)
       <223> n = A, T, C or G
      <400> 500
tccagcggnc cgccgggcng gtcatctata aaaggaaaag tgatggcatc tatatcataa
                                                                           60
atctcaagan gacctgggag aagcttctgc tggcagctcg tgcaattgtt gccattgaaa
                                                                          120
accetgetga tgtcagtgtt atatecteca ngaataetgg ccaaangget gtgetgaant ttgetgetge actggaacca etceaattge tggeegette acteetggaa eetteactaa
                                                                          180
                                                                          240
ccagatcagg caacetteeg ggaccaeggn tinttgtggt tactgacece aaggetgace
                                                                          300
accaacetnt caeggaggea tittatgita acctaectae cattgegetg tgtaacaeaa
                                                                          360
gattettete tgeetatgtg gaeattggea tteeatgeaa caaccaaggg gageteacte
                                                                          420
aatgggtttg atgtggtgga tetgeteggg naagtetgeg catgeetgge accattteeg
                                                                          480
tgaacaccat ggagggatgc ctgattttac cttggccgga cacnctangg cgaattcacc
                                                                          540
acttggngcc gtatantgga tccactcgga ccaacttggg naaaatggca naatnttccg
                                                                          600
gggaaatgat ccctccaan
                                                                          619
      <210> 501
      <211> 605
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (605)
      <223> n = A,T,C or G
      <400> 501
accacactga gatagtgttt gccaggacct cccctcagca gaagctcatc attgtggaag
                                                                           60
gctgccaaag acagggtgct atcgtggctg tgactggtga cggtgtgaat gactctccag
                                                                         120
ctttgaagaa agcaaacatt ggggttgcta tggggattgc tggctcagat gtgtccaagc
                                                                         180
aagctgctga catgattett ctggatgaca actttgcctc aattgtgact ggagtagagg
                                                                         240
aaggtegtet gatetttgat aacttgaaaa aatecattge ttatacetta accagtaaca
                                                                         300
ttccccgaga tcaccccgtt cctgatattt attattgcaa acattccact accactgggg
                                                                         360
```

```
actgtcacca tectetgeat tgacttggge actgacatgg gtnetgeeat etcetggett
                                                                       420
atgagcaggc tggagggcat catgaanaaa cagcccaaaa tccaaacaga caacttgtga
                                                                       480
atgancnggt gatcacatgg ctatggcaga atggatgatc nagncctggg aggttcttac
                                                                       540
ttacttggaa tctgntgaaa cggttcttcc aatacctntt ggcctccatg gntggaanac
                                                                       600
cctga
                                                                       605
      <210> 502
      <211> 627
      <212> DNA
      <213> Homo sapiens
      <221> misc feature
      <222> (1)...(627)
      <223> n = A, T, C or G
      <400> 502
acatettget ggaaaatget geeeaggget etggagaegg tggetgeeeg ggeteeette
                                                                        60
actgtccagg tcctgaaaga ctcttgttca tgaactgtct cttcacaaag caagtccacc
                                                                       120
acttgctggg tttatcattc tgagggtcga aaactttctc acaaagtctc agtccagtct
                                                                       180
cttgccttag ctgttgtaaa taggctctca tcacttcatc ttctgtttgt ttgcaggttt
                                                                       240
ggcataaatt gcgttaagtg gaaaaccagg ctctccagga atgggaaaat taagtgattc
                                                                       300
ccagcgtata catttette teacettgge ttttggaatt geacttttge agtttettea
                                                                       360
nacattcaga aatgtagaga gttatatata tcaangnoot atcaacttca ttottaattt
                                                                       420
cataagtttt gaaaaaaaca ttggcccttg aagtaataaa tngntttatt cccaaaatct
                                                                       480
ggatentitg genetetngg ggeangneee tigaaatgae tittgatagg gaacaangee
                                                                       540
ctggtttcca nnagnttggg ttcnggaccn taaaaaaaa gggaanccgg nttttggngg
                                                                       600
gcccggttta acccaagggc cggancn
                                                                       627
      <210> 503
      <211> 629
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(629)
      <223> n = A, T, C or G
      <400> 503
ggtacattag tagagetete caateacagg cagaegecag tgteetatga ecagggggea
                                                                        60
aatatggcca aacagattgg agcagctact tatatcgaat gctcagcttt acagtcggaa
                                                                       120
aatagcgtca gagacatttt tcacgttgcc accttggcat gtgtaaataa gacaaataaa
                                                                       180
aacgttaage ggaacaaate acagagagee acaaagegga tttcacacat geetageaga
                                                                       240
ccagaactet eggeagttge taeggaetta egaaaggaea aagegaagag etgeaetgtg
                                                                       300
atgigaatci ticattatci ttaatgaaga caaaggaatc tagigtaaaa aacaacagca
                                                                       360
aacaaaaagg tgaagtctaa atgaagtgca cagccaaagt catgtatcca gaggcttang
                                                                       420
aggegtttga gangatacte atetttttgg aatnetgeet taggttegge atgtanacea
                                                                       480
agtgatgaga agtgaatcca tggaagagtt ttaatgtgac ttggaaaata tgccaaaaaa
                                                                       540
tgagagatee aataaettna ggaaaataag ggggateeaa tneetneeeg geggeeetta
                                                                       600
ggggaattca aacactnggg gcggtatan
                                                                       629
```

<210> 504

```
<211> 462
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(462)
      <223> n = A, T, C \text{ or } G
      <400> 504
acgogggago tgagtgtccc gcggggcccg aagcgtttac tttgaaaaaa ttagagtgtt
                                                                            60
caaagcaggc ccgagccgcc tggataccgc agctaggaat aatggaatag gaccgcggtt
                                                                           120
ctattttgtt ggttttcgga actgaggcca tgattaagag ggacggccgg gggcattcgt
                                                                           180
attgcgccgc tagaggttaa attcttgqac cggcgcaaqa cgqaccaqaq cgaaaqcatt
                                                                           240
tgccaagaat gttttcatta atcaagaacg aaagtcggag gttcgaagac gatcagatac
                                                                           300
cgtcgtagtt ccgaccataa acgatgcccg accggcgatg cggcggcgtt attccatgac
                                                                           360
cegnegggea getteeggga aaccaaagte tttgggttee ngggggagta tnggtgeaaa
                                                                           420
aaaaaaaaa aaaaaaaa gtcctnggnc gcgaccccct aa
                                                                           462
      <210> 505
      <211> 628
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(628)
      \langle 223 \rangle n = A,T,C or G
      <400> 505
actititit tittititi titigggggag gitatatggg titaatagit titttaatit
                                                                            60
atttaggggg aatgatggtt gtctttggat atactacagc gatggctatt gaggagtatc
                                                                           120
ctgaggcatg ggggtcaggg gttgaggtct tggtgagtgt tttagtgggg ttagcgatgg
                                                                           180
aggtaggatt ggtgctgtgg gtgaaagant atgatggggt ggtggttgtg gtaaacttta atagtgtagg aagctgaata atttatgaag gagagggtc agggttgatt cgggaggatc
                                                                           240
                                                                           300
ctattggtgc gggggctttg tatgattatg ggcgttgatt agtantaatt actggttgaa
                                                                           360
cattgtttgt tggtgtatat attgnaattg agattgctcg ggggaatang ttatgtgatt
                                                                           420
aggaataggg ttangatgag tgggaagaaa aaaagaaagg aantaaaagt ttaattattc
                                                                           480
cctttttggg ttgaagngat natggaaggg gaaaatttgg gccttgaaat tgtttaagta
                                                                           540
atacttttct aataaggtaa gtctagaaga atagggcngg ttttggtctt aaaaaggcta
                                                                           600
aaaggggatt ggcggggtgg atccnccc
                                                                           628
      <210> 506
      <211> 612
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(612)
      <223> n = A, T, C or G
      <400> 506
```

```
acggtagaac tgctattatt catcctatgt gggtaattga ggagtatgct aagattttgc
                                                                             60
cgtagetggg tttggtttaa tecaceteaa etgeetgeta tgatggataa gattgagaga
                                                                            120
9t999gagaa ggcttacgtt cagtgaggga gagatttggt atatgattga gatgggggct
                                                                            180
agtttttgtc atgtgagaag aagcaggccg gatgtcagag gggtgccttg ggtaacctct gggactcaga agtgaaaggg ggctattcct agttttattg ctatagccat tatgattatt
                                                                            240
                                                                            300
aatgatgagt attgattggt agtattggtt atggttcatt gccggagaag tatattgttg
                                                                            360
aagaggatag ctattagaag gattatggat gccgttgctt gcctgaagaa atacttgatg
                                                                            420
gcagcttctg tggaaccaag gtttatttt ttggntagaa ctggaataaa acctacatgt
                                                                            480
ttatttctan gccactcagg taaaaaatca tgcnaactta accettgata atqtqcctcc
                                                                            540
aaaatgtaaa aaaataacgg ttggcccggg ataatcccgt ncttggccga ccccctaggn
                                                                            600
aattccccc tq
                                                                            612
       <210> 507
      <211> 632
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(632)
      <223> n = A, T, C or G
      <400> 507
ggtactacgt tgtagcccac ttccactatg tcctatcaat aggagctgta tttgccatca
                                                                             60
taggaggett catteactga tttcccctat tctcaggeta caccctagac caaacctacg
                                                                            120
ccaaaatcca tttcactatc atattcatcg gcgtaaatct aactttcttc ccacaacact ttctcggcct atccggaatg ccccgacgtt actcggacta ccccgatgca tacaccacat
                                                                            180
                                                                            240
gaaacatcct atcatctgta ggctcattca tttctctaac agcagtaata ttaataattt
                                                                            300
tcatgatttg agaagccttc gcttcgaagc gaaaagtcct aatagtagaa gaaccctcca
                                                                            360
taaacctgga gtgactatat ggatgccccc caccctacca cacattcgaa gaacccgtat
                                                                            420
acataaaatc tagacaaaaa aggaaggaat cgaacccccc aaactgggtt nagccaaccc
                                                                            480
catgggcttc acgacttttt tataaaaaaa aaaaaaaaa aaaagtcctg gcccgggngg
                                                                            540
eggteanggn gaaatteaac nactgggngg eggtetaang ggteeaacte gggneeaace
                                                                            600
tgggggaaaa tgggaaagtg gttcctgggg aa
                                                                            632
      <210> 508
      <211> 336
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (336)
      <223> n = A,T,C or G
      <400> 508
eggteeteta atgetgeten eeeggeegea ntgtgattgg atatettgea gaattegeee
                                                                             60
ttagcgtggt cgccgggccg aggtacaact tccaaaaagg agacattgga gaanaaccaa
                                                                            120
gctgggtcta taaggaattg cacatgagat ggcacacata tttatgctgt ctgaaggnca
                                                                            180
cgatcatgtt accatatcaa gctgaaaatg tcaccactat ctggagattt cgaccgtgtt
                                                                            240
ttcctctctg aatctgttat gaacacnttg gttggctgga ttcantaata aatatgtaag
                                                                            300
gcctttcttt tcaaaaaaaa aaaaaaaaa aaaagt
                                                                            336
```

```
<210> 509
      <211> 624
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(624)
      \langle 223 \rangle n = A,T,C or G
      <400> 509
ggtacttttt tttttttt tttttttta tagatacaat tggcttttat ttgtgattca
                                                                         60
tgagtcaggg cagtttccat tctgcaaaat atagtgatag ctcctactgg gcaatacaac
                                                                        120
agtanaacag tgggttttgt aaaatgggaa tccaggaaca gaagaatata aataaattga
                                                                        180
tttaaataaa ctgattggtt aatttcagaa tacttcatat tactttttc taagagttaa
                                                                        240
agcagaaagg actiticitae tgigetgaet canacageet ggaeteteat gittitagga
                                                                        300
aaattttgct gttctgggat ctacctgctt cctcatgttt cagtgngagt atatggcatt
                                                                        360
taacatgact ggctccattc tggagtccca ggctgtccct aaatgagaag ttgactaaac
                                                                        420
ataaggnatt aacactactg ncaggtacca tcattttggc ttncatcatt catanggtat
                                                                        480
gatgnecene naateatace tttatttgag tttttgneat teenneceaa aaaaaaaatt
                                                                        540
ttgaanttta ccaaaggntg catgccacnt ttaaagggtt anaaaatcnc cccncenggn
                                                                        600
actaatnttg ggccatengn nggc
                                                                        624
      <210> 510
      <211> 619
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (619)
      <223> n = A,T,C or G
      <400> 510
acggatgcta cttgtccaat gatggtaaaa gggtagctta ctggttgtcc tccgattcag
                                                                        60
gttagaatga ggaggtctgc ggctaggagt caataaagtg attggcttag tgggcgaaat
                                                                       120
attatgettt gitgittgga tatatggagg atggggatta ttgctaggat gaggatggat
                                                                        180
agtaataggg caaggacgcc tcctagtttg ttagggacgg atcggagaat tgtgtaggcc
                                                                       240
aataggaaat atcattcggg cttgatgtgg ggaggggtgt ttaangggtt ggctagggta
                                                                       300
taattgtctg ggtcccctaa gaggtctggt gagaatagtg ttaatgtcat taangagaga
                                                                       360
aagaaaaaa ataagcccga gggcgtcttt gattgtgtan taaaggtgga angtgatttt
                                                                       420
atcngaatgg gaagtgattn ctaaggggtt ggtttgatcc ctttcgtgcc aaaataagaa
                                                                       480
gnggaatgct gctagggctc cataatgaag gcaanataaa atgaaagnaa aaaatctgta
                                                                       540
aggnnggact gctactaata ncctcccaaa tcttgaacaa gntttnccaa ttntggatgg
                                                                       600
nggtataant tnaattcnn
                                                                       619
      <210> 511
      <211> 634
      <212> DNA
      <213> Homo sapiens
      <220>
     <221> misc_feature
```

```
<222> (1)...(634)
      \langle 223 \rangle n = A,T,C or G
      <400> 511
cgaggacgcg gggagatggc ctagaagcaa tgatagccat cactgagaac acctagcacc
                                                                         60
caatcttggt tcctaatacc attctcccat caaaggaacc agagatcctt ggagaaatgg
                                                                        120
ttaaggaatg aggcaggaaa tatacaagat aagcctggag catcttatag ctctagaaag
                                                                        180
taagaaagta cctgcctatt ttagaatcct agagaacatt tcattgtaag aaactagccc
                                                                        240
attatttaag tgtccacagt attittcatt tcagtggtcc aagatgcgaa ggtttccaga
                                                                        300
cacaatettg tietetaata etgeteeagg tgggatatea attetgteee catgatttge aatgatgata ecegtteeet ttaatgaaac attetttnea aatgteacat ettetgaaac
                                                                        360
                                                                        420
tgngaggnga tccaattcaa gcatatctgg gntactttcc aaatcntctt agataatctt
                                                                        480
gaaccttcgt aaaagaactg gctaattaan ccanggccct gnaggaaatt ccccttttcc
                                                                        540
tcattggcag anancetgca ttaaantntt aagggttgnn ttnccnccan aaactgtgtg
                                                                        600
gtttgnaggc aaaaacggt cttgggcatt ancc
                                                                        634
      <210> 512
      <211> 623
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(623)
      <223> n = A, T, C \text{ or } G
      <400> 512
ggtacgcggg cattgttcat gactttaaca agaaacttac agcctattta gatcttaacc
                                                                         60
tggataagtg ctatgtgatc cctctgaaca cttccattgt tatgccaccc agaaacctac
                                                                        120
tggagttact tattaacatc aaggetggaa cetatttgee teagteetat etgatteatg
                                                                        180
agcacatggt tattactgat cgcattgaaa acattgatca cctgggtttc tttatttatc
                                                                        240
gactgtgtca tgacaaggaa acttacaaac tgcaacgcag agaaactatt aaaggtattc
                                                                        300
agaaacgtga agccagcaat tgtttcgcaa ttcggcattt tgaaaacaaa tttgccgtgg
                                                                        360
aaactttaat tigictigaa cagicaagaa aaacattati gaggaaaati aatatcacag
                                                                        420
cataccccc cctttacatt ttgngcagng gatatttttt aaagcttctt tnatgtaagt
                                                                        480
540
                                                                        600
cgtctttggg accaacncgg gcc
                                                                        623
      <210> 513
      <211> 623
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1) ... (623)
      <223> n = A,T,C or G
      <400> 513
actgccctct ccagatcagc agttcaggag agcacaggag gcaaaacaca gattgctggg
                                                                         60
cttattggtg ccatcatcgt gctgattgtc gttctagcca ttggatttct cctggcacct
                                                                        120
ctacaaaagt ccgtcctggc agctttagca ttgggaaact taaagggaat gctgatgcag
                                                                        180
```

```
tttgctgaaa taggcagatt gtggcgaaag gacaaatatg attgtttaat ttggatcatg
                                                                          240
acctteatet teaceattgt cetgggaete gggttaggee tggeagetag tgtggeattt
                                                                          300
caactgctaa ccatcgtgtt caggacccaa tttccaaaat gcagcacgct ggctaatatt
                                                                          360
ggaagaacca acatctataa gaataaaaaa gattattatg atatgtatga gccagaagga
                                                                          420
gtgaaaattt cagatgtcca tettetatet aetttgenaa cattggntte tttaggengg
                                                                          480
aacttatcga tgctggtngg ctttagtnca ctttgnaatt tacgcaagcc ccacaactt
                                                                          540
tgaggaaatc ccaaactgcn aancangntt nttcagtggt acccaanggt ttttttcct
                                                                          600
tggcccgacn ccctangnga atn
                                                                          623
       <210> 514
       <211> 627
       <212> DNA
       <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (627)
      <223> n = A, T, C or G
       <400> 514
ggtactcatg cccgactgtc taccaggcac acagactttg aggagagggc gtatgtcgtc
                                                                           60
ttgatccgca tcaatgatgg gggtcggcca cccttggaag gcattgtttc tttaccagtt
                                                                          120
acattetgea gttgtgtgga aggaagttgt tteeggeeag eaggteacea gaetgggata
                                                                          180
cccactgtgg gcatggcagt tggtatactg ctgaccaccc ttctggtgat tggtataatt
                                                                          240
ttagcagttg tgtttatccg cataaagaag gataaaggca aagataatgt tgaaagtgct caagcatctg aagtcaaacc tctgagaagc tgaatttgaa aaggaatgtt tgaatttata
                                                                          300
                                                                          360
tagcaagtgc tatttcagca acaaccatct catcctatta cttttcatct aacgtgcatt
                                                                          420
ataatttttt aaacagatat tooctottgt cotttaatat ttgotaaata tttottttt
                                                                          480
gangnggagt cttgctctgt cgnccaagct ggantacctg ncccggccgg ccgtcaaagg
                                                                          540
cgaattcaac aactggcggc cgtactaatg gatcgacctc ggaccaactt ggggaacatg
                                                                          600
gcanactngt tcctgngnaa aggatcc
                                                                          627
      <210> 515
      <211> 605
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (605)
      <223> n = A, T, C or G
      <400> 515
accattggtg gccaattgat ttgatggtaa gggagggatc gttgacctcg tctgttatgt
                                                                          60
aaaggatgcc gtanggatgg gagggcgatg aggactagga tgatggcggg caggatagtt
                                                                          120
cagacggttt ctatttcctg agcgtctgag atgttagtat tagttaagtt ttgttgtgag
                                                                          180
tgttaggaaa agggcataca ggactaggaa gcagataagg aaaatgatta tgagggccgt
                                                                          240
gatcatgaaa ggtgataagc tcttctatga taggggaaag taancgtctt gtanacctac
                                                                          300
ttgcgctgca tgtgccatcc cgccgtaccc taacccgtgc aaaggtagca taatcacttg
                                                                          360
tteettaatt aagggaeetg tatgaatgge tteaceaggg tteaactgte tettaetttt
                                                                          420
aaccagtgaa attgacctgc ccctgaanag gcggcnttac acaccagacg agaaaacctt
                                                                          480
tggagcttaa ttattatcca acatacctng ccggaccccc taaggcgaat tccaccactt
                                                                          540
geggegteta tggatecact eggaceactt ggggaaaagg etactgteet ggnaatgttt
                                                                         600
```

```
cctcn
                                                                           605
       <210> 516
       <211> 464
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1)...(464)
       <223> n = A,T,C \text{ or } G
       <400> 516
ggtacaacta atccgtgaca aattaccaga ttaattttac tttatttctt caggcctggg
                                                                            60
gtttttcgat gagttcaaat ttgggatctt caaatttgaa ggtgggaaat gtattcatgt
                                                                           120
ctgcattacc aaacatttgc ttgagcttaa aaagctccct ctccagctct tgctgatact
                                                                           180
etgaactage atcaacaggt cetecagatg tetgtegett agattigtat tetetaatet
                                                                           240
tgtccacaaa gagtttctgt ataggatcaa gttccttatt aaatgccact gctgtaacac
                                                                           300
caatgtteet eegeaaatgg actgagaegg etgacegaat gacagaggag aacetgaaga
                                                                           360
geetetgaag aateatgetg attettgeae teagtecega getgneaaag cettegeege
                                                                           420
caccacctte gntctaccce egegtacetg eceggeggge gete
                                                                           464
      <210> 517
      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (611)
      <223> n = A,T,C or G
      <400> 517
acceggagea eggagatete geeggettta egtteacete ggtgtetgea geaceeteeg
                                                                            60
ettectetee taggegaega gacceagtgg ctagaagtte accatgteta tteteaagat
                                                                           120
ccatgccagg gagatettig actetegeag gaateccact gttgaggttg atetetteac
                                                                           180
ctcaaaaggt ctcttcagag ctgctgtgcc cagtggtgct tcaactggta tctatgaggc
                                                                           240
cctagagete egggacaatg ataagacteg ctatatgggg aagggtgtet caaaggetgt
                                                                           300
tgagcacatc aataaaacta ttgcgcctgc cctggttagc aagaaactga acgtcacaga
                                                                           360
acaagagaag attgacaaac tgatgatcga gatggatgga acagaaaata aatctaagtt
                                                                           420
tggtgcgaac gccattctgg gggtgtcctt tgccgtctgc naaactggtg ccgttgagaa gggggtcccc tgtccttggc cggacacnct aaggcgaatt ccacacactg cggccgtact
                                                                           480
                                                                           540
atggatcgac tcggnaccaa cttgggtaat atgggcatac tggtnctggn gaaatgtttc
                                                                           600
cctccaatcc a
                                                                           611
      <210> 518
      <211> 435
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (435)
```

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<223> n = A, T, C or G<400> 518 cgaggtactt tnttttttt tttttttt tttttaagagg aaaacccggt aatgatgtcg 60 gggttgaggg ataggaggag aatgggggat aggtgtatga acatgagggt gttttctcgt 120 gtgaatgagg gttttatgtt gttaatgtgg tgggtgagtg agccccattg tgttgtggta 180 aatatgtaga gggagtatag ggctgtgact agtatgttga gtcctgtaag taggagagtg 240 atatttgatc aggagaacgt ggttactagc acagagagtt ctcccagtag gttaatagtg 300 gggggtaagg cgaggttagc gaggcttgct agaagtcatc aaaaagctat tagtgggagt 360 agagtttgaa gtccttgaaa gaggattatg atgccactgt gaatgccttc ctagtttgag 420 tttgctagcc cgcgt 435 <210> 519 <211> 407 <212> DNA <213> Homo sapiens <220> <221> misc\_feature <222> (1)...(407)  $\langle 223 \rangle$  n = A,T,C or G <400> 519 actitntttt tittttttt titttttt ncagettige aaccatacte ecceggaae 60 ccaaagactt tggtttcccg gaagctgccc ggcgggtcat gggaataacg ccgccgcatc 120 geeggtegge ategtttatg gteggaacta enaeggtntn tgategtntt enaaceteeg 180 actttcgttc ttgattaatg aaaacattct tggcaaatgc tttcgctctg gtccgtnttg 240 cgccggtcca anaatttcac ctctagcggc gcaatacnaa tgcccccggc cgtccctctt 300 aatcatggcc tcagttccga aaaccaacaa aataaaaccg cggtcctatt ccattatgcc 360 tagetgeggt atceaggegg teeceggtac etnggeegng accaege 407 <210> 520 <211> 613 <212> DNA <213> Homo sapiens <220> <221> misc feature <222> (1)...(613) <223> n = A, T, C or G<400> 520 accttctggg gcatacaaca tggcagcagg gcctcgggaa gaggggtagg aggaccgagc 60 agcattetet gtagaggaag acaggaaagg agaceetett ggcacacatt tatggagggt 120 tgtccctgaa gagaagggca ggtgggagag gttccctgtt acttaagaga aggcaccagt 180 ggcaaagagc acaatgaaga ggatgatgat aaaaacaatc acgcagataa ggacaatcat 240 cttcacgttc ttccaccaga attttcgagc caccttctgc gatgtcgtct tgaagtgctc 300 agatgtggct tccagatcct ctgtcttgtt gcggagatgt tccaagtttt ccccccgggc 360 caggateege tecacattet gggteataat attettaaet ecetecacet caetttgeag 420 gttccgcaca cgatcatttc cttcaccttc actggcttnc tncatgtctc aaagcaccca 480 geoggeagta agtgaatege ctateggntt cttecaggng ggectanttn anttetggtg 540 gtcaactttc cccgcgtact tgggcggacc ccctaagggg aattcactgg cggccgtctt 600 tggatccacc cqn 613

```
<210> 521
       <211> 606
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (606)
       <223> n = A,T,C or G
       <400> 521
actgcagtaa aagctttaac aggtggaatt gcccacttat tcaaacagaa taaggttgtt
catgtcaatg gatatggaaa gataactggc aaaaatcaag tcactgctac gaaagctgat
                                                                            60
                                                                           120
ggcggcactc aggttattga tacaaagaac attettatag ccacgggttc agaagttact
                                                                           180
cettitectg gaatcacgat agatgaagat acaatagtgi catciacagg tgetitatet
                                                                           240
ttaaaaaaag ttccagaaaa gatggttgtt attggtgcag gagtaatagg tgtagaattg
                                                                           300
ggttcagttt ggcaaagact tggtgcagat gtgacagcag ttgaattttt angtcatgta
ggtggagttg gaattgatat ggagatatet aaaaacttte aacgcateet teaaaaacag
                                                                           360
                                                                           420
gggtttaaat ttaaattgaa tacaanggta ctggtgctcc aagaagcana tggaaaaatt
                                                                           480
gatgttctat tgaanctctt ttgngggaaa gctgaantnt acttggatgn cctnggccgn
                                                                           540
acnonotagg caateeneca etggngeent ntttggteen eetggteeaa etgggnnann
                                                                           600
nggctn
                                                                           606
       <210> 522
       <211> 617
       <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (617)
      <223> n = A,T,C or G
      <400> 522
acttgcgctt actttgtagc cttcatcagg gtttgctgaa gatggcggta tataggctga
                                                                           60
gcaagaggtg gtgaggttga tcggggttta tcgattacag aacaggctcc tctagaggga
                                                                          120
tatgaagcac cgccaggtcc tttgagtttt aagctgtggc tcgtagtgtt ctggcgagca
                                                                          180
gttttgttga tttaactgtt gaggtttagg gctaagcata gtggggtatc taatcccagt ttgggtctta gctattgtgt gttcagatat gttaaagcca ctttcgtagt ctattttgtg
                                                                          240
                                                                          300
tcaactggag ttttttacaa ctcangtgag ttttagcttt attggggagg gggtgatcta
                                                                          360
aaacactett tacgeegget tetattgact tgggttaate gtgtgaceeg eggtggetgg
                                                                          420
cacgaaattg accaaccetg gggttagtat aacttaatta aactttentt attgetnaag
                                                                          480
gtaatcctgg tggttnccct gggggngtng ntaggctaaa cgtttgaacc tcattctgcg
                                                                          540
geetganett ggeeetttta tegggggatt aaaaggggae tnettgaaen gggngettet
                                                                          600
tggnaaatta taaaaca
                                                                          617
      <210> 523
      <211> 608
      <212> DNA
      <213> Homo sapiens
      <220>
```

```
<221> misc_feature
       <222> (1)...(608)
       <223> n = A, T, C or G
      <400> 523
cgaggtactt ttttttttt ttttttttt ttttggaana agtaagcctt tatttccttg
                                                                           60
ttttgcaaat aaaactggct aagttggttg ctttttggtg attaagtcaa aganaccaaa
                                                                          120
teceatatee tegteegact ceteegacte tteettgget teaacettan etggggetge
                                                                          180
agcagcagca ggagcagctg tggtggtagc aaccacaggg gcagcancca caaaggcaga
                                                                          240
tggatcaacc aanaaggeet tgacetttte aacaagtggg aaggngtaat eegteteeca
                                                                          300
aacaaagtca ggactcgttt gtctcttcaa aaaaaaaaag cganggctcg catttggtcc
                                                                          360
cctttggaca ttttgcaact cttcaatggg gttncattgg tnggtgatgg tataaacctt
                                                                          420
tgangnacct gcccggccgg ccgtcaaang gcaaattcac ccactggcgg ccgttctatg gatccnaccc ggncccaact tgggtaatat ggcanactgt tcctggggga aatgtntccc
                                                                          480
                                                                          540
tnaaattccc acaaanacaa nccgaaccta aangtaancn gggggccaag agggcnaccn
                                                                          600
ccttattq
                                                                          608
      <210> 524
      <211> 398
      <212> DNA
      <213> Homo sapiens
      <400> 524
ggtacaggat cctctaaaga gaccgcctgg ctgggtgctc aaaccacatg ggccgaccca
                                                                           60
aaagacgtca aaaccaagag ctgctcagga ggcactaaat gttgacggtc ttggccggct
                                                                          120
tcacatcete aattteagea gacageeage ggtaagtgeg atgacgeege ageaceteaa
                                                                          180
tggccttgag ttccagtggt gttgcctgaa taccaaggtc ttctaagcca ggcaggtgag
                                                                          240
gcaatttcat gtctgtgatg tgcatccgct ccactttatc ccttgttatc cagggctcaa
                                                                          300
atgggcttat ttcaaagact cttgctaccc atcgataggc aaaaagcggc aaggggaatg
                                                                          360
ggaggaacaa tctgtgagcc acaacaaaga tgtacctg
                                                                          398
      <210> 525
      <211> 607
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(607)
      \langle 223 \rangle n = A,T,C or G
      <400> 525
actiftcctg ttggcccgag tggagactgg tgttctcaaa cccggtatgg tggtcacctt
                                                                           60
tgctccagtc aacgttacaa cggaagtaaa atctgtcgaa atgcaccatg aagctttgag
                                                                          120
tgaagetett eetggggaca atgtgggett caatgteaag aatgtgtetg teaaggatgt
                                                                          180
tegtegtege aacegitget ggigacagea aaaaigacee accaaiggaa geageigget
                                                                          240
tcactgetca ggtgattate etgaaceate caggecaaat aagegeegge tatgeceetg
                                                                          300
tattggattg ccacacggct cacattgcat gcaagtttgc tgagctgaag gaaaagattg
                                                                          360
ategeegtte tggtaaaaag etggaaaatg geeetaaatt ettgaaatet ggtgatgetg
                                                                          420
ccattggtga tatgggtcct ggcaagccca tgtgtgtttg agagcttctc aaactattca
                                                                          480
ccttgggtcc tttgctgtcg tgatatgaaa aaacagtgcg ggggtgtatc aaacatggac
                                                                          540
aaaagnttnt tgacttgcag gtaccaattt nccaaaacta aaaggtnaan aaatttncca
                                                                          600
aaccgcc
                                                                          607
```

```
<210> 526
      <211> 624
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(624)
      \langle 223 \rangle n = A,T,C or G
cgaggtacgc gggggaagct ctgtttggtg ctttggatcc atttccatcg gtccttacag
                                                                         60
Ccgctcgtca gactccagca gccaagatgg tgaagcagat cgagagcaag actgcttttc
                                                                        120
aggaageett ggacgetgea ggtgataaac ttgtagtagt tgacttetea gecacgtggt
                                                                        180
gtgggccttg caaaatgatc aagcctttct ttcattccct ctctgaaaag tattccaacg
                                                                        240
tgatattcct tgaagtagat gtggatgact gtcaggatgt tgcttcagag tgtgaagtca
                                                                        300
aatgcatgcc aacattccag ttttttaaga agggacaaaa ggtgggtgaa ttttctggag
                                                                        360
ccaataagga aaagcttgaa gccaccatta atgaattagt ctaatcatgt tttctgaaaa
                                                                        420
tataaccage ccattggcta tttaaaactt gtaatttttt taatttacca aaatntaaaa
                                                                        480
tntgaagach taacccagtt ghcatctgcg tgacaathaa acattaatgc tacactttta
                                                                        540
aaaaaaaaa aaaaaaaaa gtcctgccng cggccctcaa aggggaattc cacacctggg
                                                                        600
ggccgtcttt nggncccacc cqnn
                                                                        624
      <210> 527
      <211> 611
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
      <222> (1)...(611)
      <223> n = A,T,C or G
      <400> 527
acagagtgac actgaacaga tcacaaagca cgagaaacat tagttctctc cctccccagc
                                                                        60
gteteetteg teteeetggt ttteegatgt ceacagagtg agattgteee taagtaactg
                                                                        120
catgatcaga gtgctgtctt tataagactc ttcattcagc gtatccaatt cagcaattgc
                                                                        180
ttcatcaaat gccgtttttg ccaggctaca ggccttttca ggagagttta gaatctcata
                                                                        240
gtaaaagact gagaaattta gtgccagacc aagacgaatt gggtgtgtag gctgcatttc
                                                                        300
tttcttacta atttcaaatg cttcctggta agcctgctgg gagttcgaca cagtggtttg
                                                                       360
tttgttgctc cagatgccac ttcagaaaga tcctaaaata atctcctttc attttcaagt
                                                                        420
agaacacctt actttctggt tgtgtagcat tgggaataaa atatttgtcc acagcttcag
                                                                        480
aacatcattg cagatgtcct gcagtctggc thtatctttt acggnacctc ggccgggaca
                                                                        540
ccctanggcg aattecacae etggcggccg tetantggae ngetnggcea ettgggnana
                                                                        600
tggctactgt t
                                                                        611
      <210> 528
      <211> 615
      <212> DNA
      <213> Homo sapiens
      <220>
```

```
<221> misc feature
      <222> (1)...(615)
      <223> n = A.T.C or G
      <400> 528
ggtacttttt tttttttt tttttttga gacggagtct tgttcagctg cccaggctgg
agtgcagtgg ctcgatcttc gctcactgca accaccgtct cctgggttca agcgattctc
                                                                     120
ctgtctcage ctcccaagta gctgggatta caggccacca ccatcatgcc cggctaattt
                                                                     180
ttgtatattg gtagagacgg agtttcacta tgttgggcag gctggtcttg aactcctcac
                                                                     240
ctcaggtgat ccgcccgtct tggcctccca aagtgctagg attacaggcg taagccacca
                                                                     300
tgcctggcca gatgatgtat ttaaatatca taccaaactc tgtgtattta tataaagaaa
                                                                     360
420
cccttaccat tgntgcatat tgtgcagtat aaaacacaca cttattngga catganaaaa
                                                                     480
ccgnaagaaa gncccgggta aactggactt tgccgccttt aaaaataaaa tcnaataagn
                                                                     540
gccttgaggc cctttttcaa tgcaattttt taacccggac ctgccnggng gcggtaaggg
                                                                     600
naatccancn ctggn
                                                                     615
      <210> 529
      <211> 352
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(352)
      <223> n = A,T,C or G
     <400> 529
cgaggtactt tnttttttt tttttttt tttttgggaa aagtcatgga ggccatgggg
                                                                      60
ttggcttgaa accagctttg gggggttcga ttccttcctt ttttgtctan attttatgta
                                                                     120
tacgggttct tcgaatgtgt ggtagggtgg ggggcatcca tatagtcact ccaggtttat
                                                                     180
ggagggttct tctactatta ggacttttcg cttcgaagcg aaggcttctc aaatcatgaa
                                                                     240
aattattaat attactgctg ttagagaaat gaatgagcct acagatgata ggatgtttca
                                                                     300
tgtggtgtat gcatcggggt agtccgagta acgtcggggc attcccccgc gt
                                                                     352
     <210> 530
     <211> 769
     <212> DNA
     <213> Homo sapiens
     <220>
     <221> misc_feature
     <222> (1) ... (769)
     \langle 223 \rangle n = A,T,C or G
     <400> 530
ggtactgcat agattaaaga aataaactgc agtaaagcca ctcgtaagga atgaacgcca
                                                                      60
ttgccaatga taatcctctg cacataggtg gaaatagcaa agaagtatag ttgcttcaga
                                                                     120
acaggtaata accaaaatga taaacaccag aaataggaag ccaaacatgt aatacatctg
                                                                     180
gtgtgaccaa atactattca gaatgaagaa aagttgtata aagatgcagc caaagggcaa
                                                                     240
aateceteee atgataatae caggeaaggg ettegtgtag aacgaetgtt caggaatetg
                                                                     300
acngtggaat ctgattggtt cgaactgggt gttcaatggc atcttcttaa aaccaangta
                                                                     360
tgcaccaata aacgtcnnag gcacagatat gtanaccaaa gggccaatat ggcaancagt
                                                                     420
```

```
gtncccaaaa gaaatactgt tgganatcct ctncccagag gtcagattnt tattaagaat
                                                                       480
enceeegegt etttttttg ttttttttt getecaettt nnggtaaann acntttnttt
                                                                        540
aaaaatgttt aantetantt ectaattee atnttetttn getnennnne tgetggnggn
                                                                       600
etttaaggga anteneennt ggnggegten atganeeact tgnnactggn tantagenae
                                                                       660
gttcgggang ttcccncntt ctaatatccg gnagtaannc ggctttgncn cctantggnn
                                                                       720
engetttteg aacntgeetn anannnteeg gaggtgtatn ttettetnn
                                                                       769
      <210> 531
      <211> 777
      <212> DNA
      <213> Homo sapiens
      <220>
      <22I> misc_feature
      <222> (1)...(777)
      <223> n = A, T, C or G
      <400> 531
60
ageggaagag gtgatttatt atatggttgt tacaetegge cacaaataaa cacagaaata
                                                                       120
gtccanaatg tcacaggtcc aggacagagg accaacatgg gcattttgtt tatgagcaag
                                                                       180
gtgggtetna naggtgateg gegateagag ggegatgaag ttetagatee attgagacaa
                                                                       240
getetagaca gtageatgea gteccaeaac ttgtetecaa agatteaggt ttaeteaegt
                                                                       300
catccagcan agaatggaaa gtcaaatttc ctgaattgct atgtgtctgg gtttcatcca
                                                                       360
tccgacattg aagttgactt actgaanaat ggagagagaa ttgaaaaant nggacattca
                                                                       420
taactgnntt teancaagga etggtettte tatetettgg nettnntttt tettntattt
                                                                       480
tttttntaca tngggcctta ctttaaaaac atacntttcc nnnttacncn tggatgccaa
                                                                       540
tngatttena nanattteen agnggaatee tttngttatt nttaaaantt gggatetntn
                                                                       600
gccancactt ggctaantnt taccnncttt nggaatngtc ntatgntcat tnttggaaat
                                                                       660
tnececetn anguntttet ttnngngnta aaaattntta atunttaaat tuttttena
                                                                       720
anattnntca aatactaana ntnntnnggg nttanannaa tnntgtanat gggnnng
                                                                       777
      <210> 532
      <211> 764
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (764)
      <223> n = A, T, C or G
      <400> 532
actttacaag atagattgta taagaagcca aataatgaaa gcctagaaaa aactaattta
                                                                        60
tacttatctg aaggttacaa attagacttt taaattttct ttgtagttgg tggtgtttga
                                                                       120
gggttggcta gaaatgaaag cctggatttt gtgccatgtt tgtaatatag tttgttcctt
                                                                       180
gatcaaataa tcagagaaaa gaaacttaaa gatctttgtc tgtgaagaag aaaattatct
                                                                       240
ccctagttca atctgtagtg aaataagact acagaaggca ttgttttttc ctttttattt
                                                                       300
tntgnattat atattttttt taaatatgtt ttattgtctt ctctaagcaa aaagttctta
                                                                       360
ataaacatag tatttctctc tgcgtcctat ttcattagtg aagacatagt tcacctaaaa tggcatnctg ctctgaatct agctttttat aaatggctat gtttttgatg atatgtcaca
                                                                       420
                                                                       480
ttcaaaatgg cctaattaaa tgtgttaaat gnaatggcac tcttataacc ttaaaataac
                                                                       540
canaattaac cctccaaaaa aanaaaaaaa aaaaaggcct tggccgacnc ntangngant
                                                                       600
```

```
caccnetgng genteatgga enettggeea ettgngaann nggtnangnt eeggganatt
                                                                           660
tececatnee aatteanegg acatagnnae enggeenaag ngnneeantg nngnnnnet
                                                                           720
tnnngaacng geeetnaacn eeeggggngg tngtteneee tene
                                                                           764
      <210> 533
      <211> 773
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(773)
      \langle 223 \rangle n = A,T,C or G
      <400> 533
cgaggtactt ttttttttt ttttacagat acaattggct tttatttgtg attcatgagt
                                                                           60
cagggcagtt tccattctgc aaaatatagt gatagctcct actgggcaat acaacagtag
                                                                          120
aacagtgggt tttgtaaaat gggaatccag gaacagaaga atataaataa attgatttaa
                                                                          180
ataaactgat tggttaattt cagaatactt catattactt ttttctaaga gttaaagcag
                                                                          240
aaaggacttt cttactgtgc tgactcagac agcctggact ctcatgtttt taggaaaatt
                                                                          300
ttgtctgttc tgggatctac ctgcttcctc atgttcagtg tgagtatatg gcatttagca
                                                                          360
tgactggtcc attctggagt caccaggctt gcacctaaat gagagttgac taancatagg
                                                                          420
enttaacact actgcagtac catcatting acttcatcat catanggtat gatgnentct
                                                                          480
aatnttncat tatttgagtt tggcattcag ccacgagaga atattgcctt tgacaatgnt
                                                                          540
gcatgcaact ttaaaggttt tagatnegee neenggnaet atttnngaaa tegggggtee
                                                                          600
ecenanting agittinact ggengacenn tgaenaceat taagganigt taganinece
                                                                          660
ttgaaccccc tttacaccnt ttgnatttcc cggcntaacc ccgggcnnta agggatccnt
                                                                          720
tggcntnngg cccngcnatn gaagnachtt ngannacgcc tccncaccan nng
                                                                          773
      <210> 534
      <211> 730
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (730)
      <223> n = A, T, C or G
      <400> 534
acacagacaa atttatgcga ccagggcaga ggctgtagat gattcatatt tccaattggg
                                                                           60
agggaggact cgcttggtct tataatatcg agccaaacgg tgaatccggc tctctattag aatcagacgg aatttagcat ccttatcctt tctgttcctc tcaagatgct ttcgaacagc
                                                                          120
                                                                          180
aactgctttc ttaattaaat ggtagagatc ttcaggaaga tcaggagcaa gtcccttaga
                                                                          240
cttaagaatt cttaaaattt tattgcctgt cacaaaacgt acaaattgac caggctgttg
                                                                          300
acggctgcct ccacgtcggt ggaataattc tgacgaatct gggagctcat ggttggttgg
                                                                          360
caagaaggag ctaccacaaa aacngtgctg caggtccaga agcaggagat ggccgaaaaa
                                                                          420
tgtcccgaag ttcaaccgag aggaaatcga ggcggccgag cttgaagaag tcccgattgt
                                                                          480
tegteaacet gtgaacagaa caaceeegga cegenantge ceggtnetgg ceggacacet
                                                                          540
angggaaten accetgngge gtetangace acttggeeaa etggganntg gaaatnteeg
                                                                          600
ggaaagnten teaateecaa ttaeegaena agaaetggge naagggtene atatgggene
                                                                          660
gccttnnnga nctnccctta annccccgga gggtgntggn tctcntctan nntnnngtgg
                                                                          720
nggnnaanag
                                                                          730
```

```
<210> 535
       <211> 809
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (809)
       \langle 223 \rangle n = A,T,C or G
       <400> 535
gcgtggtcng cggccgaggt accaactgca gagccaggaa aactttgaag ccttcatgaa
                                                                              60
ggcaateggt ctgceggaag ageteateea gaaggggaag gatgtcaagg gggtgtegga
                                                                             120
aatcgtgcag aatgggaagc acttcaagtt caccatcacc gctgggtcca aagtgatcca
                                                                             180
aaacgaattc acggtggggg aggaatgtga gctggagaca atgacagggg ataaagtcaa
                                                                             240
gacagtggtt canttggaag gtgacaataa actggtgaca actttcaaaa acatcaagtc
                                                                             300
tgtgaccgaa ctcaacggng acataatcac caataccatg acattgggtg acattgtctt
                                                                             360
caagagaatc agcangagaa tttaaacaag tctgcatttc atattattt antgntgtaa
                                                                             420
aattaatgta attaaagtga actttgttta aaaaaagann nntnntntaa atanaaaaaa
                                                                             480
gtneetgeet ggeggeeggt caaaggeeaa ttecagenae tngnggeent actagtgate
                                                                             540
nactogiona actigogiaa nniggoatac tiginoingg taaaintato cotoncaton
                                                                             600
ccaaattenn ccgagettaa atntaaactg gggeetatag gnneacteet tttggttgen etgeenttnn acgaactteg neeetttat antgeeece ganagggtng tttggettte
                                                                             660
                                                                             720
ntnntatatt ctctctcc ttgnnggttt ttanggtngg tcatntgggn tctntanttt
                                                                             780
agcttngaan ntantngntn tttnttnnt
                                                                             809
       <210> 536
       <211> 755
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (755)
      <223> n = A,T,C or G
      <400> 536
acttttttt tttttttt tttttttt atgaggaaaa cccggtaatg atgtcggggt
                                                                              60
tgagggatag gaggagaatg ggggataggt gtatgaacat gagggtgttt tctcgtgtga
                                                                             120
180
tgtagaggga gtatagggct gtgactagta tgttgagtcc tgtaagtagg agagtgatat
                                                                             240
ttgatcagga gaacgtggtt actagcacag agagttctcc cagtaggtta atagtggggg
                                                                             300
gtaaggegag gttagegagg ettgetagaa gteateaaaa agetattant gggagtanag tttgaagtee ttgagagagg attatgatge nacttgtaat genttegant ttgagtttge
                                                                            360
                                                                            420
tagengaata nnatgaggat gtanteengg gecaatatna aaataeteee egtnaaettn
                                                                             480
aggggttnga taaaatgctg tctacconng actttgccgn acaccttagg caattcanca ctggngccgt ctnanggncc cacttggncc acnttggnga acatggcnnc ngtcntngga
                                                                            540
                                                                             600
aatgtttent caatteeene ttenacegan tantgnaaen ggggeanaag encecatatn
                                                                             660
gtccctccct tctngaactt nnccnttaaa tncccccgga gggttnatgg ctttctctnc
                                                                             720
taananntnt tnngnggnnt tcnataanna taann
                                                                             755
      <210> 537
```

```
<211> 794
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc_feature
       <222> (1) ... (794)
       \langle 223 \rangle n = A,T,C or G
       <400> 537
cgaggtacga aagggacaag agaaataagg cctacttcac aaagcgcctt cccccgtaaa
                                                                                    60
tgatatcatc tcaacttagt attataccca cacccaccca agaacagggt ttgttaaaaa
                                                                                   120
aaaaaaaaaa aaaaaaaaaa aaaaaagtac cttgactttg ttcacagcat gtagggtgat
                                                                                   180
gagcactcac aattgttgac taaaatgctg cttttaaaac ataggaaagt agaatggttg
                                                                                   240
agtgcaaatc catagcacaa gataaattga gctagttaag gcaaatcagg taaaatagtc
                                                                                   300
atgattctat gtaatgtaaa ccagaaaaaa taaatgttca tgatttcaag atgttatatt
                                                                                   360
aaagaaaaac tttaaaaatt attatatatt tatagcaaaa gttatcttaa atatgaattc
                                                                                   420
tgttgtaatt taatgctttt gaatacagag atntaaatga agtattatct gtaaaaatgt
                                                                                   480
atattagagt tgtgatacag agtatatttc attcanccat nttcatacta ataatatgga tttaaanata tcctataaat tcnaattcaa nanaaannnt gntananaan aanggnctgn
                                                                                   540
                                                                                   600
cggcggcgca nggcaattca acaatgnggc gtctanggac nactggtcca cttgggaana ggcaacttnc tgggaatgat ccttcattcc canntaccna gctanttaac nggggcaaag
                                                                                   660
                                                                                   720
ggecenntta tgggnntnge ntntnnaant tgecettaaa acceeggngg gtgntggnte
                                                                                   780
tttnnntttn ngnt
                                                                                   794
       <210> 538
       <211> 766
       <212> DNA
       <213> Homo sapiens
       <220>
       <221> misc feature
       <222> (1) ... (766)
       <223> n = A,T,C or G
       <400> 538
ggtacgcggg ggaaggcctt cctttttcgt ctgggctgcc aacatgccat ccagactgag
                                                                                   60
gaagaccegg aaacttaggg gecaegtgag ecaeggeeae ggeegeatag geaageaceg gaageacece ggeggeegeg gtaatgetgg tggtetgeat caecacegga teaacttega
                                                                                  120
                                                                                  180
caaataccac ccaggctact ttgggaaagt tggtatgaag cattaccact taaagaggaa
                                                                                  240
ccagagette tgeccaactg teaacettga caaattgtgg aetttggtea gtgaacagae
                                                                                  300
acgggtgaat gctgctaaaa acaagactgg ggctgtccca tcattgatgt ggtgcgatcg
                                                                                  360
gctactacaa agttctggga aagggaaagc tccaaagcaa nctgtcatcg tgaaggccaa
                                                                                  420
atcttcacag aagagctgag gagaaaaata agantgttgg ggggcctgtg tctggtgctt gaagcccatt ganggagttt aattaatgct actcttttga aaaaaanann aananaaaa
                                                                                  480
                                                                                  540
gacctgeceg geggengtaa ggeaatteac enttgngeeg tetaaggace actggecaan
                                                                                  600
tgggaanang genaanntee tgggaatngt tenteaatte eccaattaac caanaangna
                                                                                  660
acnngggcca nnnggcaccc ttatggntcc ctncctttng gaactngcct tttaatccnc
                                                                                  720
engagggtnt tgctccttnt ntttntgnnt ggggtaatna aaagtn
                                                                                  766
       <210> 539
       <211> 789
       <212> DNA
```

```
<213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (789)
      <223> n = A,T,C or G
      <400> 539
accattggtg gccaattgat ttgatggtaa gggagggatc gttgacctcg tctgttatgt
                                                                           60
aaaggatgcg tagggatggg agggcgatga ggactaggat gatggcgggc aggatagttc
                                                                          120
agacggtttc tatttcctga gcgtctgaga tgttagtatt agttagtttt gttgtgagtg
                                                                          180
ttaggaaaag gcatacagga ctaggaagca gataaggaaa atgattatga gggcgtgatc
                                                                          240
atgaaaggtg ataagctctt ctatgatagg ggaagtagcg tcttgtagac ctacttgcgc
                                                                          300
tgcatgtgcc cccgcgtact tgactttctt ttntatttnt tttattnttt ttgactactt
                                                                          360
agaattttca caattctaat aagattgttc caagtctctc atgtgcaagc tttaaaggat
                                                                          420
gactettgee atttatgtae eteggnegeg accaegetaa gggcaaatte agcaeacttg
                                                                          480
cggncgttct aagtggatcc nagetcggtc caaccitgcg tatcatggca tactggtccc
                                                                          540
tngtgaaatg tatcccttac aatcncacac atcnancccg aanctaaann taaanctggg
                                                                          600
gccaataata ctactncata atgctcnctn ctgccnttca ncnggaacnt gtgcncttnt
                                                                          660
tatnatggca acneggaagn gtggttggee tteeteteta aaaentgnng gntngttgga
                                                                          720
aggganetet aggnnneggt ccaattggan negaaattnt agetnintae naaanaintt
                                                                          780
tttttcncg
                                                                          789
      <210> 540
      <211> 747
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1) ... (747)
      <223> n = A, T, C or G
      <400> 540
acttttaagg gcataataag ggttaacatt ctaggcagta taaacacacc ccataatgca
                                                                          60
agtaataggt aatctagaga tgtggacttt attgctatat gggaattaca tttaaatttg agggcatttt atataaagaa aaatacagac ctataaagtt tggcatattc attaagttat
                                                                          120
                                                                         180
cttttaatat ttttttctag aaaacaggtg acatttgtat ctacgataaa aatttttata
                                                                         240
cagaacctac tgcctcaaac tgaatcccat caagaaaact agtttctatt gnattaagta
                                                                         300
actcaaaata aattatcact togaaaactt gctttccaca ctaaggtaag tcagactaga
                                                                         360
tgaacactcc agaattttta ctacagactg ttttaagtta gaagtgatgg caatttataa
                                                                         420
attgagaata teeteeetga tgeeetaaet ggeeaaacea aaatetaaga aageagtgae
                                                                         480
nectettaet atnatgaact tetgaatang gtagggacet eetggentan nnatgaaaan
                                                                         540
nectggeega ecceetaggg aateeneact gggggeetnn anggaeenan tggeeaantt
                                                                          600
gnnanngggn aangnneetg gnaatgteen caattenena atneegnena aagtaaengg
                                                                          660
gecenggggn annnnnangn ngnenneenn nnngaannng eeettnaann neeegngggg
                                                                          720
ggngggntet nnnennnnee nnngggg
                                                                          747
      <210> 541
      <211> 773
      <212> DNA
      <213> Homo sapiens
```

```
<220>
      <221> misc feature
      <222> (1) ... (773)
      <223> n = A, T, C \text{ or } G
      <400> 541
cgaggtacca tgaaatacat atatttcata aggttcagtt acaaaatgga ttgtttcaaa
                                                                         60
tggcaatttc ttacactaac ctgattatga aaaaaagaag tctgtatcat ctgcttccaa
                                                                        120
gtctgttatg tccaaatata ttttaattat gcatttattt tgctactttt ataaatatta
                                                                        180
gagatttcac cttaaattat ttttgtaact agttctagaa catgttttcc aattattatt
                                                                        240
tttctaatgg agacatataa ttgacctatg tttatgcata tatgttctct acacagtgaa
                                                                        300
actttttta aaaagaatag taaagaaaat gcggaagctc tggctctcca aggcaaagtc
                                                                        360
aaaaaaaaaa aaaaagcggg ggggaatgcg aggaacattt tattacacct cctgatttca
                                                                        420
etecttgagt ttattttete cettggttat tggttaatge tagaaactgn attetaagag
                                                                        480
agcatecttt teaggtgaen tgataattgg aagatttgat eetteegega eetgneegge
                                                                        540
ggccgtcnaa nggcnattcc anccactggc qqcqqtctaa nqqatcnact tqqnccacct
                                                                        600
ggctaactgg caacnggtcc ngggngaaat gnatccttaa atccncactc nacccgacct
                                                                        660
aangaactgg ggcaagggnc accctatggn gctcngcctt cnngaantnn ccnncttaan
                                                                        720
aaccngggn gntggnntct nnnnnannnn cnnnntgngg gnntaanaag ann
                                                                        773
      <210> 542
      <211> 770
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc_feature
      <222> (1)...(770)
      \langle 223 \rangle n = A,T,C or G
      <400> 542
cgaggtactt ttttttttt ttttttttt tttttttag aattctgaat tttattagag
                                                                         60
aatatateta aaatacaata titattaagt tatgatatat tgnetgaatg gaaatataet
                                                                        120
ctgnatcgca actctaatta taacaatttt tacagataat acttcattta tatctctgna
                                                                        180
attcaaaagt cattaaatta caacagaatt catatttaag ataactttgc tataaatata
                                                                        240
taataatttt taaagttttt ctttaatata acatcttgaa atcatgaaca tttattttt
                                                                        300
ctgggttaca ttcatagaat catgactatt ttacctgatt tgccttaact agctcaattt
                                                                        360
atcttggcta tggatttgca ctcaccattc tactttccta tgtttaaaag cacattttag
                                                                        420
tcacaattgn gagtgctcat caccctacat gctgtgacaa aggcaagggc ctgcccgggc
                                                                        480
ggccgtncaa anggcgaatt ccncaactgg cggcggtcca agtggancga ctcggaccaa
                                                                       540
ctngggaaca tggcaactgg tcccggggaa atggaaccgt acattcccca natcagccga
                                                                        600
ncttaggtaa acngggggcn aagggggcta cncataatgg nggtccnccc ttcatngaac
                                                                        660
cgngccctnn tatnatgcac cccggagggt nnttngcctc ctcntnnnnn ntcngntgtg
                                                                        720
gagggagtcc nggggggtnc cangggggna aaaantgccn ngncccggng
                                                                        770
      <210> 543
      <211> 748
      <212> DNA
      <213> Homo sapiens
      <220>
      <221> misc feature
```

<222> (1) ... (748)

## <223> n = A,T,C or G<400> 543

accgcgggat	gcccctcatt	tacataaata	ttatactagc	atttaccatc	tcacttctag	60
gaatactagt	atatcgctca	cacctcatat	cctccctact	atgcctagaa	ggaataatac	120
tatcgctgtt	cattatagct	actctcataa	ccctcaacac	ccactccctc	ttagccaata	180
ttgtgcctat	tgccatacta	gtctttgccg	ctgcgaagca	gcggtgggcc	tagecetaet	240
agtctcaatc	tccaacacat	atggcctaga	ctacgtacat	atgctaggcc	atatggtaac	300
tctatgttta	acattttgag	gaactgccaa	actgttttcc	aaagtgacta	cactatttta	360
cattcccacc	ttgaaggtcc	aatttctcga	cattctacca	acatgggtaa	tggctgcttt	420
ttatttagca	accttaatgg	gtgtgaagag	atactcaatg	ggaatttgat	tgattcccta	480
			ngccagagnt			540
naancttgnc	atttaacnng	cngatttatn	tgatntanaa	tnttntattt	ggancengee	600
tttaagnaag	nttaaaattn	ncaatnttgg	ggcttncttt	tggccatgan	naannttaat	660
nntannanna	attnnntncn	annnggcnng	tnaannannn			720
anaannactt	tttnnnnnna	cntggcgg				748

<210> 544

<211> 327 <212> DNA

<213> Homo sapiens

<220>

<221> misc\_feature <222> (1)...(327) <223> n = A,T,C or G

## <400> 544

acttttttt t						60
gggggtgcta t						120
ggacgatggg c	atgaaactg	tggtttgctc	cacagatttc	anagcattga	ccgtagtata	180
cccccggtcg t	gtagcggtg	aaagtggttt	ggtttaaacg	tccgggaatt	gcatctgttt	240
ttaagcctaa t	gtggggaca	gctcatgagt	gcaagacgtc	ttgtgatgta	attattatac	300
gaatgggggc t	tnaatcggg	agtacct				327